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STIC Database Tracking Number: 194532

TO: Ralph J Gitomer
Location: rem/3d65/3c18
Art Unit: 1655
Tuesday, July 11, 2006
Case Serial Number: 10/053482

From: Saloni Sharma
Location: Biotech-Chem Library
REM-1A64
Phone: (571)272-8601

saloni.sharma@uspto.gov

Search Notes

Examiner Gitomer,

See attached results.

If you have any questions about this search feel free to contact me at any time.

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Saloni Sharma
Technical Information Specialist
STIC Biotech/Chem Library
(571)272-8601

Access DB# 194532

SEARCH REQUEST FORM

Scientific and Technical Information Center

Requester's Full Name: R GITOMER Examiner #: 69630 Date: 7/2/06
Art Unit: 1655 Phone Number 30 _____ Serial Number: 10/053,482
Mail Box and Bldg/Room Location: _____ Results Format Preferred (circle): PAPER DISK E-MAIL
3 C18/3065

If more than one search is submitted, please prioritize searches in order of need.

Please provide a detailed statement of the search topic, and describe as specifically as possible the subject matter to be searched. Include the elected species or structures, keywords, synonyms, acronyms, and registry numbers, and combine with the concept or utility of the invention. Define any terms that may have a special meaning. Give examples or relevant citations, authors, etc, if known. Please attach a copy of the cover sheet, pertinent claims, and abstract.

Title of Invention: _____

Inventors (please provide full names): _____

Earliest Priority Filing Date: 11/2001

For Sequence Searches Only Please include all pertinent information (parent, child, divisional, or issued patent numbers) along with the appropriate serial number.

PLEASE SEARCH COMPOUNDS (XVIII) ON TOP PAGE
Z1, THEN CONTACT ME TO DISCUSS CLAIMS.

Thanks

TG

20916

9

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Type of Search	Vendors and cost where applicable
NA Sequence (#)	STN <u>✓</u>
AA Sequence (#)	Dialog
Structure (#)	Questel/Orbit
Bibliographic	Dr. Link
Litigation	Lexis/Nexis
Fulltext	Sequence Systems
Patent Family	WWW/Internet
Other	Other (specify)

Searcher: Sabrina
Searcher Phone #: _____
Searcher Location: _____
Date Searcher Picked Up: 7/15/06
Date Completed: 7/14/06
Searcher Prep & Review Time: 60 min
Clerical Prep Time: _____
Online Time: 40 min

> d his nofile

(FILE 'HOME' ENTERED AT 16:07:22 ON 11 JUL 2006)

FILE 'CAPLUS' ENTERED AT 16:07:32 ON 11 JUL 2006
E US2001-053482/APPS

FILE 'REGISTRY' ENTERED AT 16:07:57 ON 11 JUL 2006

FILE 'STNGUIDE' ENTERED AT 16:08:00 ON 11 JUL 2006

FILE 'REGISTRY' ENTERED AT 16:12:24 ON 11 JUL 2006

L1 STRUCTURE uploaded

L2 2 SEA SSS SAM L1
D SCAN

L3 46 SEA SSS FUL L1
D SCAN

FILE 'CAPLUS' ENTERED AT 16:14:23 ON 11 JUL 2006

L4 24 SEA ABB=ON PLU=ON L3

FILE 'REGISTRY' ENTERED AT 16:14:32 ON 11 JUL 2006

FILE 'STNGUIDE' ENTERED AT 16:14:34 ON 11 JUL 2006

FILE 'REGISTRY' ENTERED AT 16:15:50 ON 11 JUL 2006

L5 STRUCTURE uploaded

L6 0 SEA SUB=L3 SSS SAM L5

L7 0 SEA SSS SAM L5

FILE 'STNGUIDE' ENTERED AT 16:16:48 ON 11 JUL 2006

FILE 'REGISTRY' ENTERED AT 16:18:36 ON 11 JUL 2006
D QUE L1

FILE 'STNGUIDE' ENTERED AT 16:19:26 ON 11 JUL 2006

FILE 'CAPLUS' ENTERED AT 16:20:39 ON 11 JUL 2006

L8 1 SEA ABB=ON PLU=ON WO2002-US34972/APPS
E WO2002-US34972/APPS

L9 24 SEA ABB=ON PLU=ON (L4 OR L8)
SEL RN L8

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L10 11 SEA ABB=ON PLU=ON (50909-86-9/BI OR 524066-91-9/BI OR
524066-92-0/BI OR 524066-93-1/BI OR 524066-94-2/BI OR 524066-95
-3/BI OR 524066-96-4/BI OR 55779-48-1/BI OR 61869-41-8/BI OR
65417-16-5/BI OR 70217-82-2/BI)

L11 7 SEA ABB=ON PLU=ON L10 AND L3

L12 3 SEA SUB=L3 SSS FUL L5

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L13 1 SEA ABB=ON PLU=ON L12

FILE 'BEILSTEIN' ENTERED AT 16:23:44 ON 11 JUL 2006

L14 0 SEA SSS FUL L5

FILE 'MARPAT' ENTERED AT 16:23:59 ON 11 JUL 2006

L15 0 SEA SSS SAM L5

L16 0 SEA SSS FUL L5
D QUE L16

L17 FILE 'CAPLUS' ENTERED AT 16:25:00 ON 11 JUL 2006
15 SEA ABB=ON PLU=ON L9 NOT (PY>2001 OR AY>2001 OR PRY>2001)

FILE 'STNGUIDE' ENTERED AT 16:25:24 ON 11 JUL 2006

FILE 'CAPLUS' ENTERED AT 16:26:06 ON 11 JUL 2006
E WOOD K/AU

L18 919 SEA ABB=ON PLU=ON WOOD K?/AU
E HAWKINS E/AU

L19 264 SEA ABB=ON PLU=ON HAWKINS E?/AU
E SCURRIA M/AU

L20 6 SEA ABB=ON PLU=ON ("SCURRIA M A"/AU OR "SCURRIA MICHAEL"/AU
OR "SCURRIA MICHAEL A"/AU OR "SCURRIA MIKE"/AU)
E KLAUBERT D/AU

L21 72 SEA ABB=ON PLU=ON ("KLAUBERT D"/AU OR "KLAUBERT D H"/AU OR
"KLAUBERT D K"/AU OR "KLAUBERT DIETER"/AU OR "KLAUBERT DIETER
H"/AU OR "KLAUBERT DIETER HEINZ"/AU)

L22 12 SEA ABB=ON PLU=ON (L18 AND (L19 OR L20 OR L21)) OR (L19 AND
(L20 OR L21)) OR (L20 AND L21)

L23 4 SEA ABB=ON PLU=ON L11

L24 24 SEA ABB=ON PLU=ON (L23 OR L9)
D BIB L13

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FILE LAST UPDATED: 10 Jul 2006 (20060710/ED)

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'OBI' IS DEFAULT SEARCH FIELD FOR 'CAPLUS' FILE

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L18 919 SEA FILE=CAPLUS ABB=ON PLU=ON WOOD K?/AU
L19 264 SEA FILE=CAPLUS ABB=ON PLU=ON HAWKINS E?/AU
L20 6 SEA FILE=CAPLUS ABB=ON PLU=ON ("SCURRIA M A"/AU OR "SCURRIA
MICHAEL"/AU OR "SCURRIA MICHAEL A"/AU OR "SCURRIA MIKE"/AU)
L21 72 SEA FILE=CAPLUS ABB=ON PLU=ON ("KLAUBERT D"/AU OR "KLAUBERT
D H"/AU OR "KLAUBERT D K"/AU OR "KLAUBERT DIETER"/AU OR

L22 "KLAUBERT DIETER H"/AU OR "KLAUBERT DIETER HEINZ"/AU)
 12 SEA FILE=CAPLUS ABB=ON PLU=ON (L18 AND (L19 OR L20 OR L21))
 OR (L19 AND (L20 OR L21)) OR (L20 AND L21)

=> d ibib abs l22 tot

L22 ANSWER 1 OF 12 CAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 2006:137803 CAPLUS
 DOCUMENT NUMBER: 144:384691
 TITLE: New Bioluminogenic Substrates for Monoamine Oxidase Assays
 AUTHOR(S): Zhou, Wenhui; Valley, Michael P.; Shultz, John;
 Hawkins, Erika M.; Bernad, Laurent; Good, Troy; Good, Dave; Riss, Terry L.; Klaubert, Dieter H.; Wood, Keith V.
 CORPORATE SOURCE: Promega Biosciences Inc., San Luis Obispo, CA, 93401, USA
 SOURCE: Journal of the American Chemical Society (2006), 128(10), 3122-3123
 CODEN: JACSAT; ISSN: 0002-7863
 PUBLISHER: American Chemical Society
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 AB Novel bioluminogenic substrates were designed for probing monoamine oxidase (MAO) activity based on a simple and effective β-elimination strategy. By modifying the amino group and the central core of luciferin derivs., we have developed a series of substrates useful for assays of MAO A or B, or both. One of these substrates, exhibiting low Km values and high signal-to-background ratios with both isoenzymes, was shown to accurately measure the Ki values of known MAO inhibitors. This substrate is a key component in the development of a highly sensitive homogeneous MAO assay for high-throughput screening (HTS) of compds. in drug discovery and for monitoring MAO activity in complex biol. systems. This design strategy should be applicable to fluorogenic MAO substrates and could broaden the structural requirements of substrates for other enzyme assays.
 REFERENCE COUNT: 15 THERE ARE 15 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L22 ANSWER 2 OF 12 CAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 2005:1292638 CAPLUS
 DOCUMENT NUMBER: 144:33522
 TITLE: Substrate-binding, catalytically inactive hydrolases as carriers for the immobilization of fusion proteins
 INVENTOR(S): Darzins, Aldis; Encell, Lance; Johnson, Tonny; Klaubert, Dieter; Los, Georgyi V.; McDougall, Mark; Wood, Keith V.; Wood, Monika G.; Zimprich, Chad
 PATENT ASSIGNEE(S): USA
 SOURCE: U.S. Pat. Appl. Publ., 121 pp., Cont.-in-part of U.S. Ser. No. 768,976.
 CODEN: USXXCO
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 2
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
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US 2005272114	A1	20051208	US 2004-6031	20041206
US 2004214258	A1	20041028	US 2004-768976	20040130
US 2006024808	A1	20060202	US 2005-194110	20050729
PRIORITY APPLN. INFO.:			US 2003-444094P	P 20030131
			US 2003-474659P	P 20030530
			US 2004-768976	A2 20040130
			US 2004-592499P	P 20040730

OTHER SOURCE(S) : MARPAT 144:33522

AB Hydrolase variants that retain substrate binding, and capable of forming a covalent bond with a substrate, but lacking the catalytic activity to release the hydrolysis products are described for use in the immobilization of proteins onto surfaces carrying a substrate for the hydrolase are described. The binding of the hydrolase to substrate is more stable than that of the wild type enzyme. The catalytically inactive variant has at least two amino acid substitutions. Substrates for hydrolases comprising one or more functional groups are also provided, as well as methods of using the mutant hydrolase and the substrates of the invention. Also provided is a fusion protein capable of forming a stable bond with a substrate and cells which express the fusion protein. Development of a catalytically inactive variant of the haloalkane dehalogenase of *Rhodococcus rhodochrous* is demonstrated. Use of fusion products with fluorescent proteins and enzymes in imaging *in vivo* are demonstrated.

L22 ANSWER 3 OF 12 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2005:291435 CAPLUS
 DOCUMENT NUMBER: 143:341532
 TITLE: Homogeneous, bioluminescent protease assays: Caspase-3 as a model
 AUTHOR(S): O'Brien, Martha A.; Daily, William J.; Hesselberth, P. Eric; Moravec, Richard A.; Scurria, Michael A.; Klaubert, Dieter H.; Bulleit, Robert F.; Wood, Keith V.
 CORPORATE SOURCE: Promega Corporation, Madison, WI, USA
 SOURCE: Journal of Biomolecular Screening (2005), 10(2), 137-148
 CODEN: JBISF3; ISSN: 1087-0571
 PUBLISHER: Sage Publications
 DOCUMENT TYPE: Journal
 LANGUAGE: English

AB Using caspase-3 as a model, the authors have developed a strategy for highly sensitive, homogeneous protease assays suitable for high-throughput, automated applications. The assay uses peptide-conjugated aminoluciferin as the protease substrate and a firefly luciferase that has been molecularly evolved for increased stability. By combining the pro luminescent caspase-3 substrate, Z-DEVD-aminoluciferin, with a stabilized luciferase in a homogeneous format, the authors developed an assay that is significantly faster and more sensitive than fluorescent caspase-3 assays. The assay has a single-step format, in which protease cleavage of the substrate and luciferase oxidation of the aminoluciferin occurs simultaneously. Because these processes are coupled, they rapidly achieve steady state to maintain stable luminescence for several hours. Maximum sensitivity is attained when this steady state occurs; consequently, this coupled-enzyme system results in a very rapid assay. The homogeneous format inherently removes trace contamination by free aminoluciferin, resulting in extremely low background and yielding exceptionally high signal-to-noise ratios and excellent Z' factors. Another advantage of a luminescent format is that it avoids problems of cell autofluorescence or fluorescence interference that can be associated

with synthetic chemical and natural product libraries. This bioluminescent, homogeneous format should be widely applicable to other protease assays.
 REFERENCE COUNT: 25 THERE ARE 25 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L22 ANSWER 4 OF 12 CAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 2005:150211 CAPLUS
 TITLE: Analytical biotechnology
 AUTHOR(S): Wood, Keith V.; Klaubert, Dieter H.
 CORPORATE SOURCE: Promega Corporation, Madison, WI, 53711, USA
 SOURCE: Current Opinion in Biotechnology (2005), 16(1), 1-2
 CODEN: CUOBEB; ISSN: 0958-1669
 PUBLISHER: Elsevier Ltd.
 DOCUMENT TYPE: Journal; Editorial
 LANGUAGE: English
 AB Unavailable

L22 ANSWER 5 OF 12 CAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 2004:698252 CAPLUS
 DOCUMENT NUMBER: 141:187324
 TITLE: Methods and kits for dual enzymatic assays whereby light is quenched from luminescent reactions
 INVENTOR(S): Hawkins, Erika; Butler, Braeden; Wood, Keith V.
 PATENT ASSIGNEE(S): Promega Corporation, USA
 SOURCE: PCT Int. Appl., 77 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2004072299	A1	20040826	WO 2004-US4075	20040212
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI				
RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
AU 2004210982	A1	20040826	AU 2004-210982	20040212
CA 2515217	AA	20040826	CA 2004-2515217	20040212
US 2004224377	A1	20041111	US 2004-777461	20040212
EP 1592805	A1	20051109	EP 2004-710594	20040212
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK				
PRIORITY APPLN. INFO.:			US 2003-447065P	P 20030212
			WO 2004-US4075	W 20040212

AB The present invention relates to single and dual reporter luminescence assays utilizing reagents to quench an optical, e.g., an enzyme-mediated luminescence, reaction. In one embodiment of the invention, a reagent is added to an assay which selectively quenches a first enzyme-mediated luminescence reaction without affecting a subsequent distinct enzyme-mediated luminescent reaction(s). An assay kit containing one or more selective quench reagents, and compns. comprising the quench reagent(s), are also provided.

L22 ANSWER 6 OF 12 CAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 2004:698213 CAPLUS
 DOCUMENT NUMBER: 141:221282
 TITLE: Mutant Rhodococcus dehalogenase and functionalized chloroalkane substrates useful for covalent tethering of functional groups to proteins
 INVENTOR(S): Wood, Keith V.; Los, Georgyi V.; Bulleit, Robert F.; Klaubert, Dieter; McDougall, Mark; Zimprich, Chad
 PATENT ASSIGNEE(S): Promega Corporation, USA
 SOURCE: PCT Int. Appl., 185 pp.
 CODEN: PIIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 2
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2004072232	A2	20040826	WO 2004-US2607	20040130
WO 2004072232	C2	20041014		
WO 2004072232	A3	20050127		
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
AU 2004211584	A1	20040826	AU 2004-211584	20040130
CA 2514564	AA	20050726	CA 2004-2514564	20040130
EP 1594962	A2	20051116	EP 2004-707032	20040130
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK				
CN 1764721	A	20060426	CN 2004-80008194	20040130
PRIORITY APPLN. INFO.: US 2003-444094P P 20030131 US 2003-474659P P 20030530 WO 2004-US2607 W 20040130				

OTHER SOURCE(S): MARPAT 141:221282
 AB A mutant hydrolase optionally fused to a protein of interest is provided. Thus, Rhodococcus haloalkane dehalogenase DhaA with His-272 substituted with Phe is capable of forming a bond with a chloroalkane substrate for the corresponding nonmutant (wild-type) hydrolase which is more stable than the bond formed between the wild-type hydrolase and the substrate. The chloroalkane substrate contains a functional group which binds Ca²⁺ or K⁺, or Na⁺, is pH sensitive, is a radionuclide, is electron opaque, is a chromophore or fluorophore, is a MRI contrast agent, is a substance that fluoresces in the presence of NO, or is sensitive to reactive oxygen. Substrates for hydrolases comprising one or more functional groups are synthesized comprising TAMRA-, FAM-, and ROX.5-C14H24O4-Cl or biotin-C18H32O4-Cl, as methods of using the mutant DhaA and the substrates of the invention for cell imaging in vivo are provided. Mutant *Staphylococcus aureus* β-lactamase (blaZ)-based tethering of functional groups is also demonstrated. Also provided is a fusion protein capable of forming a stable bond with a substrate and cells which express the fusion protein.

L22 ANSWER 7 OF 12 CAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 2004:570131 CAPLUS
 DOCUMENT NUMBER: 141:119301
 TITLE: Improving the accuracy of luciferase-based assays for high throughput screening by using tolerance enhancement agents
 INVENTOR(S): Hawkins, Erika; Cali, James J.; Ho, Samuel Kin Sang; O'Brien, Martha; Somberg, Richard; Bulleit, Robert F.; Wood, Keith V.
 PATENT ASSIGNEE(S): Promega Corporation, USA
 SOURCE: PCT Int. Appl., 68 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2004059294	A2	20040715	WO 2003-US41454	20031223
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW				
RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
CA 2508072	AA	20040715	CA 2003-2508072	20031223
AU 2003300008	A1	20040722	AU 2003-300008	20031223
US 2005026171	A1	20050203	US 2003-746995	20031223
EP 1588143	A2	20051026	EP 2003-800272	20031223
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PRIORITY APPLN. INFO.:			US 2002-436173P	P 20021223
			US 2003-444264P	P 20030131
			US 2003-447334P	P 20030213
			WO 2003-US41454	W 20031223

AB The invention concerns methods and kits for improving the accuracy of luciferase-based assays for high throughput screening of compound libraries by reducing the number of false hits. A method and kit is provided for enhancing the tolerance of an assay reagent to compds. in an assay sample, the assay reagent including a luciferase enzyme. The method includes contacting the luciferase with a tolerance enhancement agent in an amount sufficient to substantially protect luciferase enzyme activity from interference of the compound and minimize interference by at least about 10% relative to an assay not having tolerance enhancement agent. Tolerance-enhancing effect of detergents on the inhibition of luciferase was studied. Minimization of false hit occurrence using tolerance enhancement agents such as detergents was demonstrated.

L22 ANSWER 8 OF 12 CAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 2004:270174 CAPLUS
 DOCUMENT NUMBER: 140:299425
 TITLE: Luminescent cytochrome P 450 assay using luciferase, luciferin derivatives and pyrophosphatase, and drug screening applications

INVENTOR(S) : Cali, James J.; Klaubert, Dieter; Daily, William; Ho, Samuel Kin Sang; Frackman, Susan; Hawkins, Erika; Wood, Keith V.

PATENT ASSIGNEE(S) : Promega Corporation, USA

SOURCE: PCT Int. Appl., 130 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2004027378	A2	20040401	WO 2003-US29078	20030916
WO 2004027378	A3	20041125		
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CA 2497560	AA	20040401	CA 2003-2497560	20030916
AU 2003267245	A1	20040408	AU 2003-267245	20030916
EP 1546162	A2	20050629	EP 2003-749715	20030916
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JP 2006508339	T2	20060309	JP 2004-537859	20030916
US 2004171099	A1	20040902	US 2003-665314	20030919
PRIORITY APPLN. INFO. :			US 2002-412254P	P 20020920
			US 2003-483309P	P 20030627
			WO 2003-US29078	W 20030916

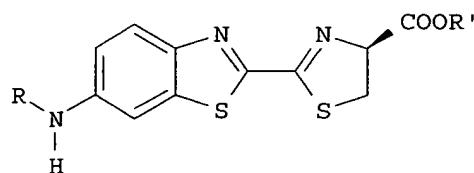
OTHER SOURCE(S) : MARPAT 140:299425

AB The present invention provides methods, compns., substrates, and kits useful for analyzing the metabolic activity in cells, tissue, and animals and for screening test compds. for their effect on cytochrome P 450 activity. In particular, a one-step and two-step methods using luminogenic mols., e.g. luciferin or coelenterazines, that are cytochrome P 450 substrates and that are also bioluminescent enzyme, e.g., luciferase, pro-substrates are provided. Upon addition of the luciferin derivative or other luminogenic mol. into a P 450 reaction, the P 450 enzyme metabolizes the mol. into a bioluminescent enzyme substrate, e.g., luciferin and/or luciferin derivative metabolite, in a P 450 reaction. The resulting metabolite(s) serves as a substrate of the bioluminescent enzyme, e.g., luciferase, in a second light-generating reaction. Luminescent cytochrome P 450 assays with low background signals and high sensitivity are disclosed and isoform selectivity is demonstrated. The present invention also provides an improved method for performing luciferase reactions which employs added pyrophosphatase to remove inorg. pyrophosphate, a luciferase inhibitor which may be present in the reaction mixture as a contaminant or may be generated during the reaction. The present method further provides a method for stabilizing and prolonging the luminescent signal in a luciferase-based assay using luciferase stabilizing agents such as reversible luciferase inhibitors.

ACCESSION NUMBER: 2003:633682 CAPLUS
 DOCUMENT NUMBER: 139:193612
 TITLE: Bioluminescent protease assay using aminoluciferin linked to peptide substrate and luciferase
 INVENTOR(S): O'Brian, Martha; Wood, Keith; Klaubert, Dieter; Daily, Bill
 PATENT ASSIGNEE(S): Promega Corporation, USA
 SOURCE: PCT Int. Appl., 55 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2003066611	A1	20030814	WO 2003-US2936	20030131
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
CA 2474695	AA	20030814	CA 2003-2474695	20030131
AU 2003216139	A1	20030902	AU 2003-216139	20030131
US 2003211560	A1	20031113	US 2003-356665	20030131
EP 1472238	A1	20041103	EP 2003-737580	20030131
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK				
JP 2005530485	T2	20051013	JP 2003-565985	20030131
US 2006121546	A1	20060608	US 2006-347054	20060203
PRIORITY APPLN. INFO.:			US 2002-353158P	P 20020201
			US 2003-356665	A3 20030131
			WO 2003-US2936	W 20030131

GI



AB A sensitive bioluminescent assay to detect proteases including caspases, trypsin and tryptase is provided. The method comprises contacting a sample suspected of having one or more caspases with a mixture comprising beetle luciferase and an aminomodified beetle aminoluciferin or a carboxyterminal protected derivative thereof, wherein the amino group of aminoluciferin or the derivative thereof is modified so as to covalently link a substrate for the caspase via a peptide bond to aminoluciferin or the carboxyterminal protected derivative thereof. If the sample comprises a caspase having a recognition site in the substrate, the substrate is

cleaved at the peptide bond that links the substrate to aminoluciferin, yielding aminoluciferin, a substrate for the luciferase, in the mixture. Luminescence is then detected. The method further comprises correlating luminescence with protease concentration or activity, i.e., increased luminescence correlates with increased protease concentration or activity.

Also

provided is a compound comprising aminoluciferin or a carboxyterminal protected derivative thereof covalently linked via a peptide bond to a protease recognition site such as a caspase recognition site, a trypsin recognition site, or a tryptase recognition site. A specific compound of the invention is a compound of formula I (R = peptide with an aspartic acid, lysine, or arginine C-terminus; R' = H, carboxy protecting group, e.g., C1-6-alkyl, Ph, benzyl ester, counterion). The invention also provides synthetic processes and intermediates disclosed herein, which are useful for preparing compds. of the invention. As described herein below, using a substrate for caspase 3 and 7 that was linked to either aminoluciferin or rhodamine-110, it was found that the limit of detection for the aminoluciferin-based substrate was 0.2 to 0.5 µU of purified caspase while that for the rhodamine-110-based substrate was 10 µU. As also described herein, it was found that the limit of detection of caspase expressing cells with the aminoluciferin-based substrate was 15 cells at 1 h while the limit of detection for the rhodamine-110-based substrate was 150 cells at 1 h.

REFERENCE COUNT: 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L22 ANSWER 10 OF 12 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2003:376823 CAPLUS

DOCUMENT NUMBER: 138:365147

TITLE: Compositions, methods and kits pertaining to luminescent compounds

INVENTOR(S): Wood, Keith; Hawkins, Erika;
Scurria, Mike; Klaubert, Dieter

PATENT ASSIGNEE(S): Promega Corporation, USA

SOURCE: PCT Int. Appl., 60 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2003040100	A1	20030515	WO 2002-US34972	20021101
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, UZ, VC, VN, YU, ZA, ZM, ZW				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
US 2003153090	A1	20030814	US 2001-53482	20011102
CA 2462506	AA	20030515	CA 2002-2462506	20021101
EP 1451155	A1	20040901	EP 2002-802815	20021101
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, SK				

JP 2005515977	T2 20050602	JP 2003-542146	20021101
PRIORITY APPLN. INFO.:		US 2001-53482	A 20011102
		WO 2002-US34972	W 20021101

OTHER SOURCE(S) : MARPAT 138:365147

AB A method of measuring the enzymic activity of a luciferase includes contacting a luminogenic protein, such as a luciferase, with a protected luminophore to form a composition; and detecting light produced from the composition

The protected luminophore provides increased stability and improved signal-to-background ratios relative to the corresponding unmodified coelenterazine.

REFERENCE COUNT: 8 THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L22 ANSWER 11 OF 12 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2003:108790 CAPLUS

DOCUMENT NUMBER: 139:129758

TITLE: Coelenterazine derivatives for improved solution solubility

AUTHOR(S) : Hawkins, Erika M.; O'Grady, Michael;
Klaubert, Dieter; Scurria, Michael;
Good, Troy; Stratford, Cathy; Flemming, Rod; Simpson,
Dan; Wood, Keith V.

CORPORATE SOURCE: Promega Corporation, Madison, WI, 53715, USA

SOURCE: Bioluminescence & Chemiluminescence: Progress & Current Applications, [Proceedings of the Symposium on Bioluminescence and Chemiluminescence], 12th, Cambridge, United Kingdom, Apr. 5-9, 2002 (2002), 149-152. Editor(s): Stanley, Philip E.; Kricka, Larry J. World Scientific Publishing Co. Pte. Ltd.: Singapore, Singapore.

CODEN: 69DPGZ; ISBN: 981-238-156-2

DOCUMENT TYPE: Conference

LANGUAGE: English

AB Intracellular luminescent techniques requiring coelenterazine, such as bioluminescence resonance energy transfer (BRET), calcium detection, and intracellular reporter measurements, must accommodate the poor stability of this substrate in physiol. buffered solns. Coelenterazine degradation leads both to loss of luminescence over time, and increased background luminescence caused by enzyme-independent oxidation (autoluminescence). Both conditions limit luminescence sensitivity by reducing the signal-to-noise ratio. Coelenterazine can be stabilized by derivatizing the enol oxygen with an acyl oxymethyl ether. This prevents spontaneous oxidation in solution while making the substrate available intracellularly upon cleavage of the blocking group by endogenous esterases. We will describe the stability of pivaloyl oxymethyl coelenterazine-h (POM coelenterazine-h), and the effect of POM coelenterazine-h on intracellular luminescence, autoluminescence, and luminescent reaction kinetics. Also, we will present the characteristics of two other coelenterazine derivs.

L22 ANSWER 12 OF 12 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2001:924094 CAPLUS

DOCUMENT NUMBER: 136:50649

TITLE: Method for increasing luminescence assay sensitivity

INVENTOR(S) : Hawkins, Erika; Centanni, John M.; Sankbeil,
Jacqueline; Wood, Keith V.

PATENT ASSIGNEE(S) : Promega Corporation, USA

SOURCE: PCT Int. Appl., 45 pp.

CODEN: PIXXD2

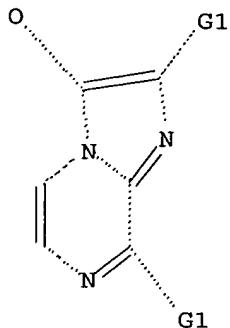
DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001096862	A2	20011220	WO 2001-US18363	20010607
WO 2001096862	A3	20020718		
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
CA 2411179	AA	20011220	CA 2001-2411179	20010607
EP 1297337	A2	20030402	EP 2001-942027	20010607
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR				
JP 2004503777	T2	20040205	JP 2002-510941	20010607
US 2004096924	A1	20040520	US 2003-692587	20031024
US 2006051827	A1	20060309	US 2004-991759	20041118
PRIORITY APPLN. INFO.:			US 2000-590884	A 20000609
			WO 2001-US18363	W 20010607
			US 2003-692587	A3 20031024

AB A method for increasing the sensitivity of a luminescent assay comprising carrying out the assay in the presence of an organic compound that reduces luminescence that is not dependent on the presence of an analyte by at least about 10 fold, and that reduces luminescence that is dependent on the presence of an analyte by less than about 7 fold.

=> d que 124
 L1 STR

Cy¹



G1 Ak,H,[@1]

Structure attributes must be viewed using STN Express query preparation.

L3 46 SEA FILE=REGISTRY SSS FUL L1
L4 24 SEA FILE=CAPLUS ABB=ON PLU=ON L3
L8 1 SEA FILE=CAPLUS ABB=ON PLU=ON WO2002-US34972/APPS
L9 24 SEA FILE=CAPLUS ABB=ON PLU=ON (L4 OR L8)
L10 11 SEA FILE=REGISTRY ABB=ON PLU=ON (50909-86-9/BI OR 524066-91-9
 /BI OR 524066-92-0/BI OR 524066-93-1/BI OR 524066-94-2/BI OR
 524066-95-3/BI OR 524066-96-4/BI OR 55779-48-1/BI OR 61869-41-8
 /BI OR 65417-16-5/BI OR 70217-82-2/BI)
L11 7 SEA FILE=REGISTRY ABB=ON PLU=ON L10 AND L3
L23 4 SEA FILE=CAPLUS ABB=ON PLU=ON L11
L24 24 SEA FILE=CAPLUS ABB=ON PLU=ON (L23 OR L9)

=> d ibib abs hitstr l24 tot

L24 ANSWER 1 OF 24 CAPLUS COPYRIGHT 2006 ACS on STN
ACCESSION NUMBER: 2005:397579 CAPLUS
DOCUMENT NUMBER: 143:419154
TITLE: Chemical studies of fish bioluminescence
AUTHOR(S): Kakoi, Hisae; Okada, Kunisuke
CORPORATE SOURCE: Faculty of Pharmacy, Meijo University, Tempaku-ku,
Nagoya, 468-8503, Japan
SOURCE: ITE Letters on Batteries, New Technologies & Medicine
(2005), 6(1), 38-45
CODEN: ILBMF9; ISSN: 1531-2046
PUBLISHER: ITE Inc.
DOCUMENT TYPE: Journal
LANGUAGE: English
AB Watasenia preluciferin (I), first isolated from the squid Watasenia
scintillans, is a compound that plays a key role in the light emitting

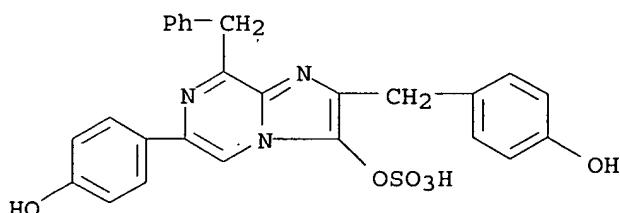
process of various bioluminescent marine organisms such as squids, shrimps, coelenterates, and fish. In the case of luminous fish, a well-known species is Myctophiformes and Stomiiformes especially the deep-sea photophores-possessing Myctophiformes fish (lantern fish), which is one of the most common and widely distributed luminous fish living in Suruga Bay and all throughout the Sea of Enshu and Kumano. Compound I was isolated either from the liver of Neoscopelus microchir (in Japanese, Sango-iwashi) or from a pair of large nasal photophores from Diaphus gigas (in Japanese, Suito-hadaka) while it was found neither in the photophores of N. microchir nor in the liver of D. gigas. On the other hand, a luciferase active toward Oplophorus luciferin (=Watasenia preluciferin) I was extracted from the flesh of D. gigas, whereas no luciferase active toward I or Cypridina luciferin was found in N. microchir. Later, a new type of bound form of I was isolated from the liver of D. gigas and the structure was established as Diaphus luciferyl β -glucopyranosiduronic acid (II) on the basis of the spectral data and chemical evidence, and by synthesis starting from I. This compound II was also detected in the liver of Diaphus watasei (in Japanese, Hadaka-iwashi) and other examined Myctophiformes fish, but not in the liver of N. microchir. It is uncertain as to which system is more favorable for the fish bioluminescence, however, as far as I is concerned, the Diaphus bioluminescent system is comparable to that of Watasenia or Oplophorus, and not to that of Cypridina as previously observed by Tsuji et al. in 1971.

IT 55779-47-0

RL: BSU (Biological study, unclassified); BIOL (Biological study)
(chemical studies of fish bioluminescence)

RN 55779-47-0 CAPLUS

CN Imidazo[1,2-a]pyrazin-3-ol, 6-(4-hydroxyphenyl)-2-[(4-hydroxyphenyl)methyl]-8-(phenylmethyl)-, 3-(hydrogen sulfate) (9CI) (CA INDEX NAME)



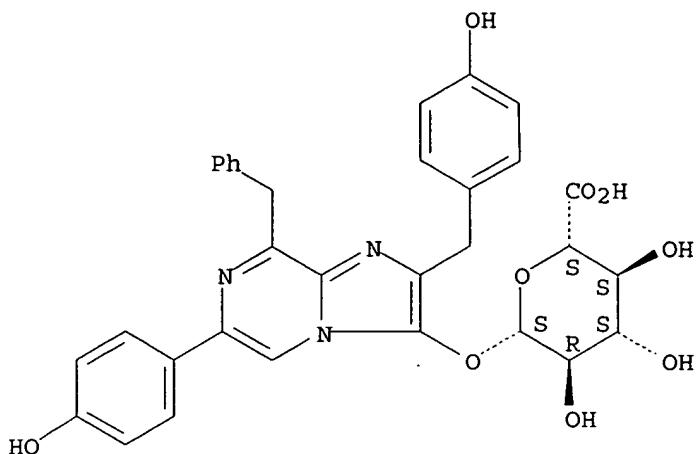
IT 107503-09-3P

RL: BSU (Biological study, unclassified); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation)
(chemical studies of fish bioluminescence)

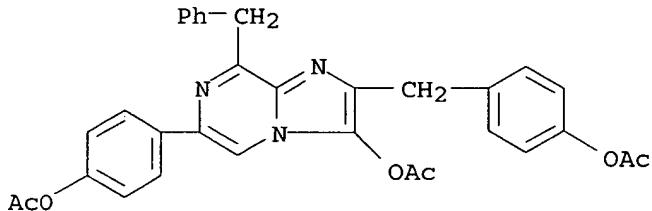
RN 107503-09-3 CAPLUS

CN β -D-Glucopyranosiduronic acid, 6-(4-hydroxyphenyl)-2-[(4-hydroxyphenyl)methyl]-8-(phenylmethyl)imidazo[1,2-a]pyrazin-3-yl (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

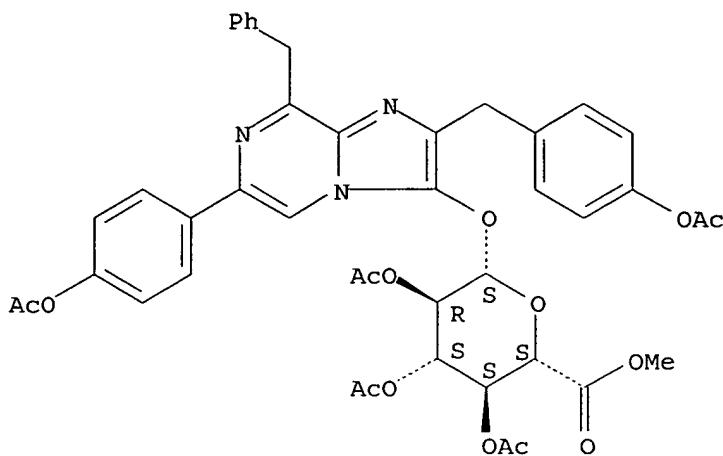


IT 65417-16-5P 107503-11-7P 867375-43-7P
 RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT
 (Reactant or reagent)
 (intermediate in preparation of luciferyl β -glucopyranosiduronic acid)
 RN 65417-16-5 CAPLUS
 CN Imidazo[1,2-a]pyrazin-3-ol, 6-[4-(acetoxy)phenyl]-2-[[4-(acetoxy)phenyl]methyl]-8-(phenylmethyl)-, acetate (ester) (9CI) (CA INDEX NAME)



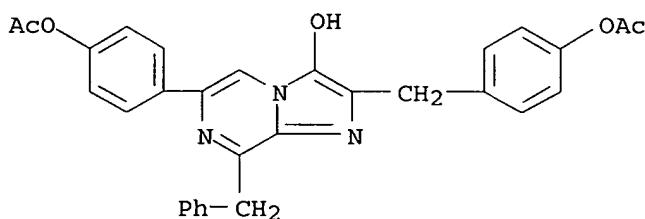
RN 107503-11-7 CAPLUS
 CN β -D-Glucopyranosiduronic acid, 6-[4-(acetoxy)phenyl]-2-[[4-(acetoxy)phenyl]methyl]-8-(phenylmethyl)imidazo[1,2-a]pyrazin-3-yl, methyl ester, 2,3,4-triacetate (9CI) (CA INDEX NAME)

Absolute stereochemistry.



RN 867375-43-7 CAPLUS

CN Imidazo[1,2-a]pyrazin-3-ol, 6-[4-(acetoxy)phenyl]-2-[(4-(acetoxy)phenyl)methyl]-8-(phenylmethyl)- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 18 THERE ARE 18 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L24 ANSWER 2 OF 24 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2004:270174 CAPLUS

DOCUMENT NUMBER: 140:299425

TITLE: Luminescent cytochrome P 450 assay using luciferase, luciferin derivatives and pyrophosphatase, and drug screening applications

INVENTOR(S): Cali, James J.; Klaubert, Dieter; Daily, William; Ho, Samuel Kin Sang; Frackman, Susan; Hawkins, Erika; Wood, Keith V.

PATENT ASSIGNEE(S): Promega Corporation, USA

SOURCE: PCT Int. Appl., 130 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2004027378	A2	20040401	WO 2003-US29078	20030916
WO 2004027378	A3	20041125		

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,

GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW			
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG			
CA 2497560	AA 20040401	CA 2003-2497560	20030916
AU 2003267245	A1 20040408	AU 2003-267245	20030916
EP 1546162	A2 20050629	EP 2003-749715	20030916
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK			
JP 2006508339	T2 20060309	JP 2004-537859	20030916
US 2004171099	A1 20040902	US 2003-665314	20030919
PRIORITY APPLN. INFO.:			
		US 2002-412254P	P 20020920
		US 2003-483309P	P 20030627
		WO 2003-US29078	W 20030916

OTHER SOURCE(S) : MARPAT 140:299425

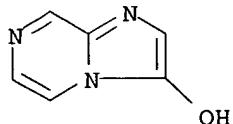
AB The present invention provides methods, compns., substrates, and kits useful for analyzing the metabolic activity in cells, tissue, and animals and for screening test compds. for their effect on cytochrome P 450 activity. In particular, a one-step and two-step methods using luminogenic mols., e.g. luciferin or coelenterazines, that are cytochrome P 450 substrates and that are also bioluminescent enzyme, e.g., luciferase, pro-substrates are provided. Upon addition of the luciferin derivative or other luminogenic mol. into a P 450 reaction, the P 450 enzyme metabolizes the mol. into a bioluminescent enzyme substrate, e.g., luciferin and/or luciferin derivative metabolite, in a P 450 reaction. The resulting metabolite(s) serves as a substrate of the bioluminescent enzyme, e.g., luciferase, in a second light-generating reaction. Luminescent cytochrome P 450 assays with low background signals and high sensitivity are disclosed and isoform selectivity is demonstrated. The present invention also provides an improved method for performing luciferase reactions which employs added pyrophosphatase to remove inorg. pyrophosphate, a luciferase inhibitor which may be present in the reaction mixture as a contaminant or may be generated during the reaction. The present method further provides a method for stabilizing and prolonging the luminescent signal in a luciferase-based assay using luciferase stabilizing agents such as reversible luciferase inhibitors.

IT 676460-49-4D, Imidazo[1,2-a]pyrazin-3-ol, derivs.

RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses)
(luminescent cytochrome P 450 assay using luciferase, luciferin derivs.
and pyrophosphatase, and drug screening applications)

RN 676460-49-4 CAPLUS

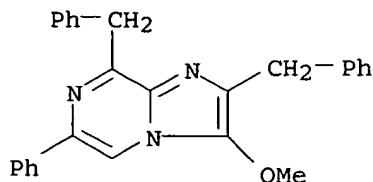
CN Imidazo[1,2-a]pyrazin-3-ol (9CI) (CA INDEX NAME)



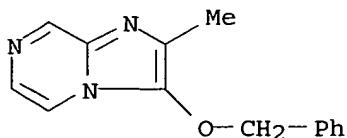
IT 676460-47-2P, Coelenterazine HH methyl ether

RL: ARG (Analytical reagent use); SPN (Synthetic preparation); ANST
(Analytical study); PREP (Preparation); USES (Uses)
(luminescent cytochrome P 450 assay using luciferase, luciferin derivs.)

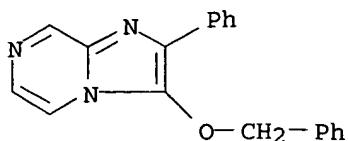
and pyrophosphatase, and drug screening applications)
 RN 676460-47-2 CAPLUS
 CN Imidazo[1,2-a]pyrazine, 3-methoxy-6-phenyl-2,8-bis(phenylmethyl)- (9CI)
 (CA INDEX NAME)



L24 ANSWER 3 OF 24 CAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 2004:40051 CAPLUS
 DOCUMENT NUMBER: 140:429758
 TITLE: Metal-ion complexation of imidazo[1,2-a]pyrazin-3(7H)-ones: continuous changes in absorption spectra of complexes depending on the Lewis acidity of the metal ion
 AUTHOR(S): Sekiguchi, Takashi; Maki, Shojiro; Niwa, Haruki; Ikeda, Hiroshi; Hirano, Takashi
 CORPORATE SOURCE: Department of Applied Physics and Chemistry, The University of Electro-Communications, Chofu, Tokyo, 182-8585, Japan
 SOURCE: Tetrahedron Letters (2004), 45(5), 1065-1069
 CODEN: TELEAY; ISSN: 0040-4039
 PUBLISHER: Elsevier Science B.V.
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 AB The spectroscopic properties of metal-ion complexes of several imidazopyrazinone derivs. with Li⁺, Mg²⁺, Ca²⁺, Ba²⁺, Sc³⁺, and La³⁺ ions were studied. The spectral characteristics and the formation consts. of the complexes changed continuously depending on the Lewis acidity of the metal ion, suggesting that the imidazopyrazinones can find application as indicators of Lewis acidity. In the case of bis-imidazopyrazinone derivs., the complexation abilities were enhanced by chelate effects.
 IT 693252-73-2P 693252-74-3P
 RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
 (formation of metal-ion complexes with imidazopyrazinones and dependence of their absorption spectra on metal-ion Lewis acidity)
 RN 693252-73-2 CAPLUS
 CN Imidazo[1,2-a]pyrazine, 2-methyl-3-(phenylmethoxy)- (9CI) (CA INDEX NAME)

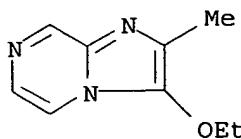


RN 693252-74-3 CAPLUS
 CN Imidazo[1,2-a]pyrazine, 2-phenyl-3-(phenylmethoxy)- (9CI) (CA INDEX NAME)

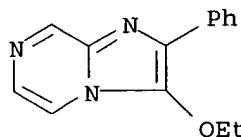


REFERENCE COUNT: 23 THERE ARE 23 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

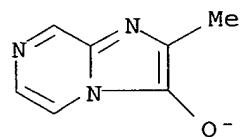
L24 ANSWER 4 OF 24 CAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 2003:1005621 CAPLUS
 DOCUMENT NUMBER: 140:181114
 TITLE: Fundamental studies on the structures and spectroscopic properties of imidazo[1,2-a]pyrazin-3(7H)-one derivatives
 AUTHOR(S): Nakai, Shunichiro; Yasui, Masanori; Nakazato, Masaki; Iwasaki, Fujiko; Maki, Shojiro; Niwa, Haruki; Ohashi, Mamoru; Hirano, Takashi
 CORPORATE SOURCE: Department of Applied Physics and Chemistry, The University of Electro-Communications, Tokyo, 182-8585, Japan
 SOURCE: Bulletin of the Chemical Society of Japan (2003), 76(12), 2361-2387
 CODEN: BCSJA8; ISSN: 0009-2673
 PUBLISHER: Chemical Society of Japan
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 OTHER SOURCE(S): CASREACT 140:181114
 AB The fundamental phys. properties of 2-Me and 2-phenylimidazo[1,2-a]pyrazin-3(7H)-one, and their N- and O-alkylated derivs. were studied by x-ray crystallog., UV/visible absorption spectroscopy, NMR, and AM1-COSMO calcns. The crystal structures of showed that the imidazo[1,2-a]pyrazin-3(7H)-one (imidazopyrazinone) π -system has a planar ring structure and a weakened carbonyl character of the C3-O10 bond, suggesting that the imidazopyrazinone π -system has the character of a zwitter-ionic resonance structure to increase the aromaticity. The data concerning the bond length alternations and the NMR chemical shifts of 1-4 also support that their imidazopyrazinone rings have small portions of aromatic character. Imidazopyrazinone derivs. 1-4 showed solvatochromism originating by H-bonding interactions with H-bond donor solvent mols.; derivs. 1 and 2 prefer to be the NH form isomers in their tautomeric equilibrium. These observations were consistently evaluated by MO calcns. The phys. properties of protonated species of 1-6 and anion species of 1 and 2 were also established. The fundamental properties of the imidazopyrazinone π -system explain the several problems of the chemi- and bioluminescence reactivities of imidazopyrazinone derivs. and of the construction of a bioluminescent supramol.
 IT 659726-97-3P 659726-99-5P
 RL: CPS (Chemical process); PEP (Physical, engineering or chemical process); PRP (Properties); RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); PROC (Process); RACT (Reactant or reagent)
 (fundamental studies on structures and spectral properties of imidazo[1,2-a]pyrazin-3(7H)-one derivs.)
 RN 659726-97-3 CAPLUS
 CN Imidazo[1,2-a]pyrazine, 3-ethoxy-2-methyl- (9CI) (CA INDEX NAME)



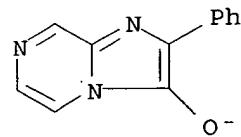
RN 659726-99-5 CAPLUS
CN Imidazo[1,2-a]pyrazine, 3-ethoxy-2-phenyl- (9CI) (CA INDEX NAME)



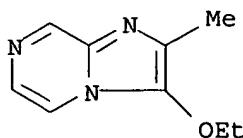
IT 659727-06-7 659727-07-8 659727-12-5
659727-13-6
RL: FMU (Formation, unclassified); PRP (Properties); FORM (Formation, nonpreparative)
(fundamental studies on structures and spectral properties of imidazo[1,2-a]pyrazin-3(7H)-one derivs.)
RN 659727-06-7 CAPLUS
CN Imidazo[1,2-a]pyrazin-3-ol, 2-methyl-, ion(1-) (9CI) (CA INDEX NAME)



RN 659727-07-8 CAPLUS
CN Imidazo[1,2-a]pyrazin-3-ol, 2-phenyl-, ion(1-) (9CI) (CA INDEX NAME)

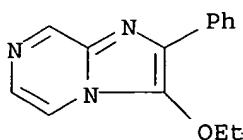


RN 659727-12-5 CAPLUS
CN Imidazo[1,2-a]pyrazine, 3-ethoxy-2-methyl-, conjugate monoacid (9CI) (CA INDEX NAME)



● H⁺

RN 659727-13-6 CAPLUS
 CN Imidazo[1,2-a]pyrazine, 3-ethoxy-2-phenyl-, conjugate monoacid (9CI) (CA INDEX NAME)



● H⁺

REFERENCE COUNT: 82 THERE ARE 82 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L24 ANSWER 5 OF 24 CAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 2003:376823 CAPLUS
 DOCUMENT NUMBER: 138:365147
 TITLE: Compositions, methods and kits pertaining to luminescent compounds
 INVENTOR(S): Wood, Keith; Hawkins, Erika; Scurria, Mike; Klaubert, Dieter
 PATENT ASSIGNEE(S): Promega Corporation, USA
 SOURCE: PCT Int. Appl., 60 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2003040100	A1	20030515	WO 2002-US34972	20021101 <--
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, UZ, VC, VN, YU, ZA, ZM, ZW				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
US 2003153090	A1	20030814	US 2001-53482	20011102

CA 2462506	AA . 20030515	CA 2002-2462506	20021101 <--
EP 1451155	A1 20040901	EP 2002-802815	20021101 <--
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, SK			
JP 2005515977	T2 20050602	JP 2003-542146	20021101 <--
PRIORITY APPLN. INFO.:		US 2001-53482	A 20011102
		WO 2002-US34972	W 20021101 <--

OTHER SOURCE(S): MARPAT 138:365147

AB A method of measuring the enzymic activity of a luciferase includes contacting a luminogenic protein, such as a luciferase, with a protected luminophore to form a composition; and detecting light produced from the composition

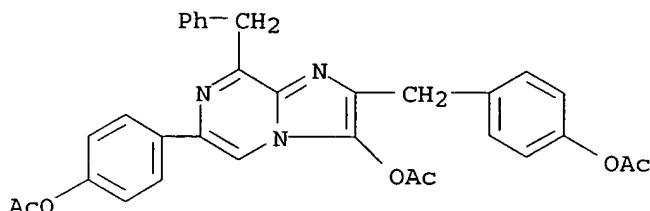
The protected luminophore provides increased stability and improved signal-to-background ratios relative to the corresponding unmodified coelenterazine.

IT 65417-16-5P 524066-91-9P 524066-92-0P
524066-93-1P 524066-94-2P 524066-95-3P
524066-96-4P

RL: ARU (Analytical role, unclassified); SPN (Synthetic preparation); ANST (Analytical study); PREP (Preparation)
(compns., methods and kits pertaining to luminescent compds.)

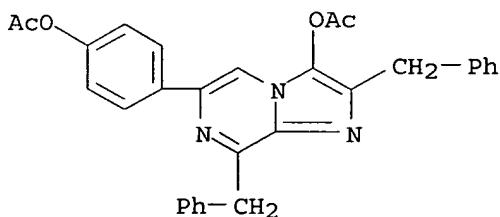
RN 65417-16-5 CAPLUS

CN Imidazo[1,2-a]pyrazin-3-ol, 6-[4-(acetyloxy)phenyl]-2-[[4-(acetyloxy)phenyl]methyl]-8-(phenylmethyl)-, acetate (ester) (9CI) (CA INDEX NAME)



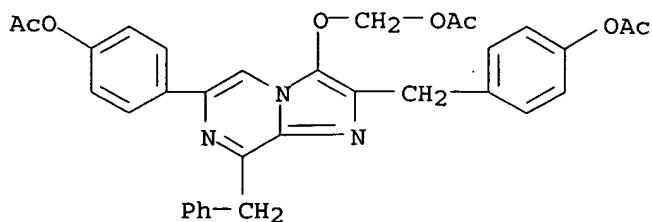
RN 524066-91-9 CAPLUS

CN Imidazo[1,2-a]pyrazin-3-ol, 6-[4-(acetyloxy)phenyl]-2,8-bis(phenylmethyl)-, acetate (ester) (9CI) (CA INDEX NAME)



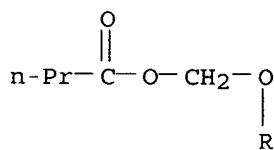
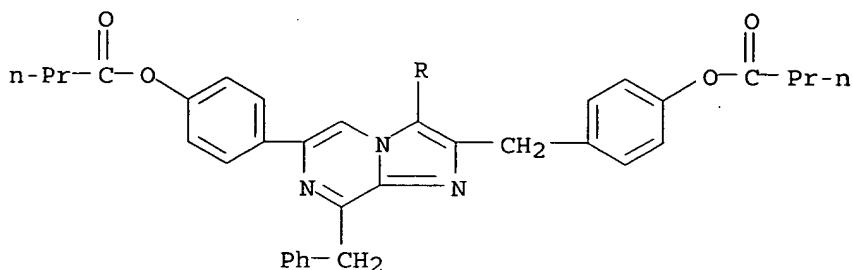
RN 524066-92-0 CAPLUS

CN Phenol, 4-[3-[(acetyloxy)methoxy]-2-[[4-(acetyloxy)phenyl]methyl]-8-(phenylmethyl)imidazo[1,2-a]pyrazin-6-yl]-, acetate (ester) (9CI) (CA INDEX NAME)



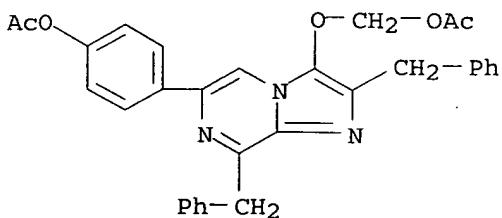
RN 524066-93-1 CAPLUS

CN Butanoic acid, 4-[3-[(1-oxobutoxy)methoxy]-2-[[4-(1-oxobutoxy)phenyl]methyl]-8-(phenylmethyl)imidazo[1,2-a]pyrazin-6-yl]phenyl ester (9CI) (CA INDEX NAME)



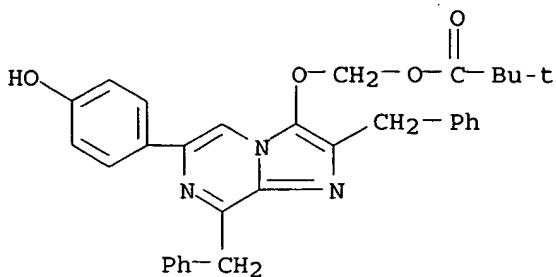
RN 524066-94-2 CAPLUS

CN Phenol, 4-[3-[(acetyloxy)methoxy]-2,8-bis(phenylmethyl)imidazo[1,2-a]pyrazin-6-yl]-, acetate (ester) (9CI) (CA INDEX NAME)



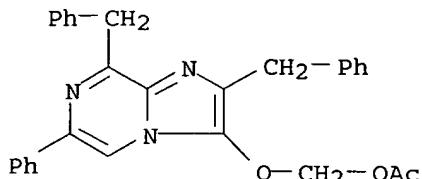
RN 524066-95-3 CAPLUS

CN Propanoic acid, 2,2-dimethyl-, [[6-(4-hydroxyphenyl)-2,8-bis(phenylmethyl)imidazo[1,2-a]pyrazin-3-yl]oxy]methyl ester (9CI) (CA INDEX NAME)



RN 524066-96-4 CAPLUS

CN Methanol, [[6-phenyl-2,8-bis(phenylmethyl)imidazo[1,2-a]pyrazin-3-yl]oxy] - , acetate (ester) (9CI) (CA INDEX NAME)



REFERENCE COUNT: 8 THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L24 ANSWER 6 OF 24 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2003:108790 CAPLUS

DOCUMENT NUMBER: 139:129758

TITLE: Coelenterazine derivatives for improved solution solubility

AUTHOR(S): Hawkins, Erika M.; O'Grady, Michael; Klaubert, Dieter; Scurria, Michael; Good, Troy; Stratford, Cathy; Flemming, Rod; Simpson, Dan; Wood, Keith V.

CORPORATE SOURCE: Promega Corporation, Madison, WI, 53715, USA

SOURCE: Bioluminescence & Chemiluminescence: Progress & Current Applications, [Proceedings of the Symposium on Bioluminescence and Chemiluminescence], 12th, Cambridge, United Kingdom, Apr. 5-9, 2002 (2002), 149-152. Editor(s): Stanley, Philip E.; Kricka, Larry J. World Scientific Publishing Co. Pte. Ltd.: Singapore, Singapore.

DOCUMENT TYPE: Conference

LANGUAGE: English

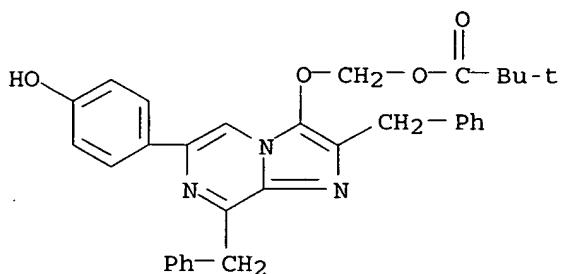
AB Intracellular luminescent techniques requiring coelenterazine, such as bioluminescence resonance energy transfer (BRET), calcium detection, and intracellular reporter measurements, must accommodate the poor stability of this substrate in physiol. buffered solns. Coelenterazine degradation leads both to loss of luminescence over time, and increased background luminescence caused by enzyme-independent oxidation (autoluminescence). Both conditions limit luminescence sensitivity by reducing the signal-to-noise ratio. Coelenterazine can be stabilized by derivatizing the enol oxygen with an acyl oxymethyl ether. This prevents spontaneous oxidation in solution while making the substrate available intracellularly upon cleavage of the blocking group by endogenous esterases. We will describe the stability of

pivaloyl oxymethyl coelenterazine-h (POM coelenterazine-h), and the effect of POM coelenterazine-h on intracellular luminescence, autoluminescence, and luminescent reaction kinetics. Also, we will present the characteristics of two other coelenterazine derivs.

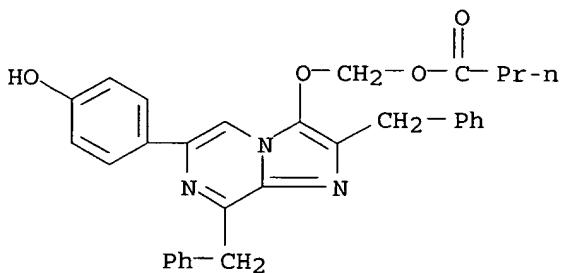
IT 524066-95-3D, diacetyl derivative 566945-96-8
 RL: BSU (Biological study, unclassified); BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)
 (coelenterazine derivs. for improved solution solubility)

RN 524066-95-3 CAPLUS

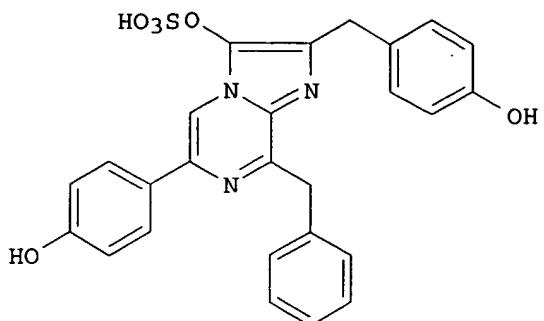
CN Propanoic acid, 2,2-dimethyl-, [[6-(4-hydroxyphenyl)-2,8-bis(phenylmethyl)imidazo[1,2-a]pyrazin-3-yl]oxy]methyl ester (9CI) (CA INDEX NAME)



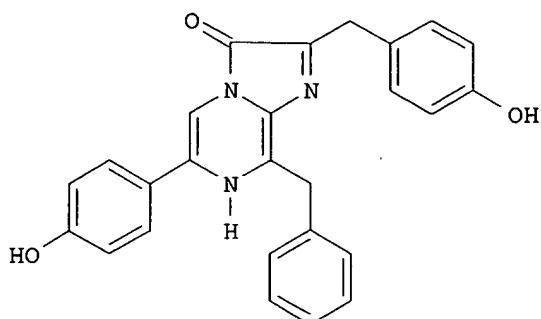
RN 566945-96-8 CAPLUS
 CN Butanoic acid, [[6-(4-hydroxyphenyl)-2,8-bis(phenylmethyl)imidazo[1,2-a]pyrazin-3-yl]oxy]methyl ester (9CI) (CA INDEX NAME)



L24 ANSWER 7 OF 24 CAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 2002:906075 CAPLUS
 DOCUMENT NUMBER: 138:153360
 TITLE: Efficient synthesis of Renilla preluciferin
 AUTHOR(S): Teranishi, Katsunori
 CORPORATE SOURCE: Faculty of Bioresources, Mie University, Tsu, Mie, 514-8507, Japan
 SOURCE: ITE Letters on Batteries, New Technologies & Medicine (2002), 3(4), 479-480
 CODEN: ILBMF9; ISSN: 1531-2046
 PUBLISHER: ITE-IBA Publication Office
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 OTHER SOURCE(S): CASREACT 138:153360
 GI

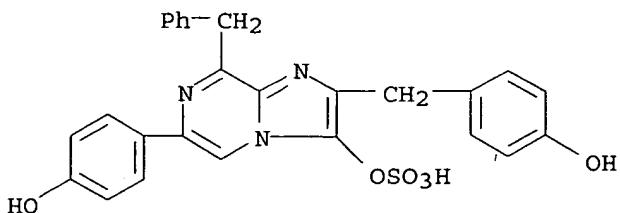


I



II

- AB Renilla luciferyl sulfate that is Renilla preluciferin I was efficiently synthesized by one-step procedure from coelenterazine (II).
- IT 55779-47-0P
RL: SPN (Synthetic preparation); PREP (Preparation)
(efficient synthesis of Renilla preluciferin via sulfation of coelenterazine)
- RN 55779-47-0 CAPLUS
- CN Imidazo[1,2-a]pyrazin-3-ol, 6-(4-hydroxyphenyl)-2-[(4-hydroxyphenyl)methyl]-, 3-(hydrogen sulfate) (9CI) (CA INDEX NAME)



REFERENCE COUNT: 9 THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L24 ANSWER 8 OF 24 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2001:851130 CAPLUS

DOCUMENT NUMBER: 135:371764

TITLE: Preparation of aminopyrazines and imidazolopyrazinones as

antioxidants

INVENTOR(S) : Marchand-Brynaert, Jacqueline; Cavalier,
Jean-Francois; Rees, Jean-Francois; De Tollenaere,
Catherine; Burton, Maggi

PATENT ASSIGNEE(S) : Universite Catholique de Louvain, Belg.

SOURCE : PCT Int. Appl., 57 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001087853	A1	20011122	WO 2001-EP5588	20010516
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
EP 1292580	A1	20030319	EP 2001-943383	20010516
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR				
US 2004034225	A1	20040219	US 2003-276398	20030728
PRIORITY APPLN. INFO.:			EP 2000-870107	A 20000517
			EP 2000-870293	A 20001212
			WO 2001-EP5588	W 20010516

OTHER SOURCE(S) : CASREACT 135:371764; MARPAT 135:371764

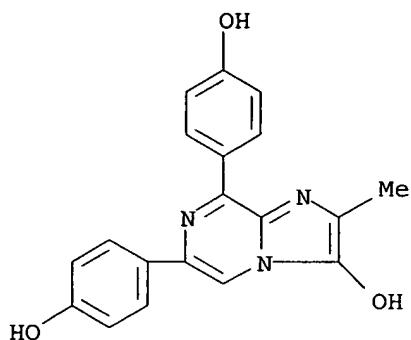
AB Antioxidants, 5 2-amino-(p-hydroxyphenyl)pyrazines and 3 (p-hydroxyphenyl)-3,7-dihydroimidazo[1,2-a]pyrazin-3-ones were prepared and claimed useful in diagnostic procedures, as food additives, polymer additives and as UV screens in cosmetics. E.g., 2-amino-3,5-dibromopyrazine was treated with p-methoxyphenylboronic acid in the presence of bis(benzonitrile)palladium dichloride and 1,4-bis(diphenylphosphino)butane in a solvent mix of EtOH, aqueous sodium carbonate and toluene to give 66% 2-amino-3,5-bis(p-methoxyphenyl)pyrazine, which was demethylated with EtSNa in DMF to give 88% 2-amino-3,5-bis(p-hydroxyphenyl)pyrazine (I). In tests on inhibition of lipid peroxidation. 2-aminopyrazines possessing 2 aryl substituents, one of them being a p-hydroxyphenyl in o- or p- position with respect to the amino group, are endowed with antioxidative properties. However, the p-hydroxyphenyl conferred more activity when located at position 5 than at position 3. The presence of p-hydroxyphenyl groups at both positions 3 and 5 as in I produced a very active compound. Analogs lacking the free phenol groups showed reduced activities. Corresponding imidazolopyrazinones combined the properties of both the imidazolopyrazinones (delay of the onset of peroxidation) and the aminopyrazines (lower rate of oxidation after onset).

IT 374588-75-7P 374588-76-8P 374588-77-9P
374588-78-0P
RL: FFD (Food or feed use); MOA (Modifier or additive use); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation); USES (Uses)
(preparation of aminopyrazines and imidazolopyrazinones as antioxidants)

RN 374588-75-7 CAPLUS

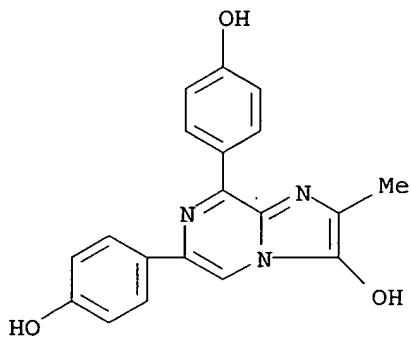
CN Imidazo[1,2-a]pyrazin-3-ol, 6,8-bis(4-hydroxyphenyl)-2-methyl- (9CI) (CA)

INDEX NAME)



RN 374588-76-8 CAPLUS

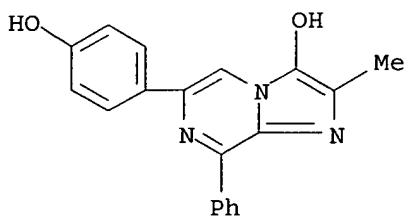
CN Imidazo[1,2-a]pyrazin-3-ol, 6,8-bis(4-hydroxyphenyl)-2-methyl-, monohydrochloride (9CI) (CA INDEX NAME)



● HCl

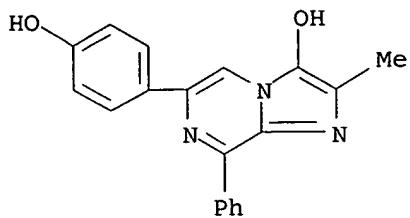
RN 374588-77-9 CAPLUS

CN Imidazo[1,2-a]pyrazin-3-ol, 6-(4-hydroxyphenyl)-2-methyl-8-phenyl- (9CI) (CA INDEX NAME)



RN 374588-78-0 CAPLUS

CN Imidazo[1,2-a]pyrazin-3-ol, 6-(4-hydroxyphenyl)-2-methyl-8-phenyl-, monohydrochloride (9CI) (CA INDEX NAME)



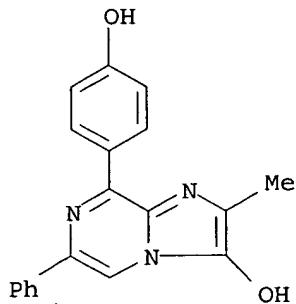
● HCl

IT 374588-79-1P 374588-80-4P 374588-85-9P
374588-86-0P 374588-87-1P

RL: SPN (Synthetic preparation); PREP (Preparation)
(preparation of aminopyrazines and imidazolopyrazinones as antioxidants)

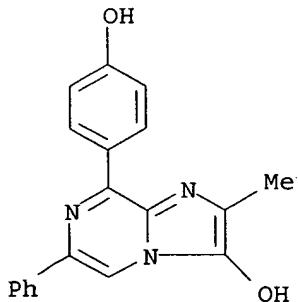
RN 374588-79-1 CAPLUS

CN Imidazo[1,2-a]pyrazin-3-ol, 8-(4-hydroxyphenyl)-2-methyl-6-phenyl- (9CI)
(CA INDEX NAME)



RN 374588-80-4 CAPLUS

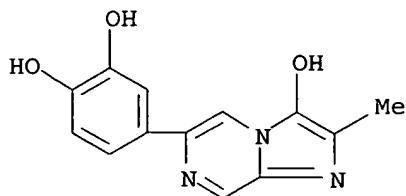
CN Imidazo[1,2-a]pyrazin-3-ol, 8-(4-hydroxyphenyl)-2-methyl-6-phenyl-,
monohydrochloride (9CI) (CA INDEX NAME)



● HCl

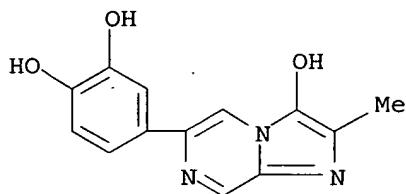
RN 374588-85-9 CAPLUS

CN 1,2-Benzenediol, 4-(3-hydroxy-2-methylimidazo[1,2-a]pyrazin-6-yl)- (9CI)
(CA INDEX NAME)



RN 374588-86-0 CAPLUS

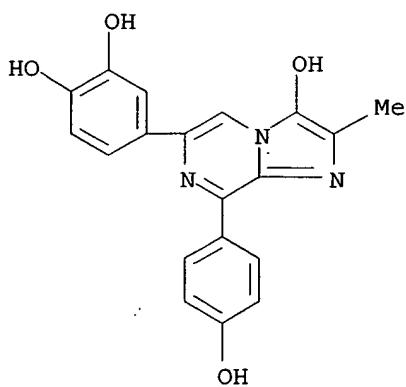
CN 1,2-Benzenediol, 4-(3-hydroxy-2-methylimidazo[1,2-a]pyrazin-6-yl)-, monohydrochloride (9CI) (CA INDEX NAME)



● HCl

RN 374588-87-1 CAPLUS

CN 1,2-Benzenediol, 4-[3-hydroxy-8-(4-hydroxyphenyl)-2-methylimidazo[1,2-a]pyrazin-6-yl]- (9CI) (CA INDEX NAME)



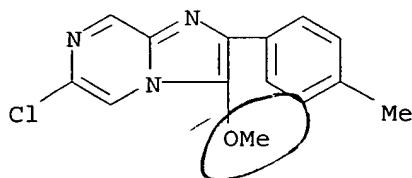
REFERENCE COUNT:

10

THERE ARE 10 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L24 ANSWER 9 OF 24 CAPLUS COPYRIGHT 2006 ACS on STN
ACCESSION NUMBER: 1998:725609 CAPLUS

DOCUMENT NUMBER: 130:104779
 TITLE: Imidazo[1,2-b]pyridazines: syntheses and interaction with central and peripheral-type (mitochondrial) benzodiazepine receptors
 AUTHOR(S): Barlin, Gordon B.
 CORPORATE SOURCE: Division of Neuroscience, John Curtin School of Medical Research, Australian National University, Canberra, ACT 2601, Australia
 SOURCE: Journal of Heterocyclic Chemistry (1998), 35(5), 1205-1217
 CODEN: JHTCAD; ISSN: 0022-152X
 PUBLISHER: HeteroCorporation
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 AB The fundamental chemical of pyridazines, the syntheses of substituted imidazo[1,2-b]pyridazines (1) (and some related compds.) and the interaction of the products with central benzodiazepine receptors (CBR) and peripheral-type (mitochondrial) benzodiazepine receptors (PBR) are described. Some of these imidazo[1,2-b]pyridazines had high selective affinity for the central benzodiazepine receptors and others had high selectivity for the peripheral-type (mitochondrial) benzodiazepine receptors. The results of structure-activity studies and mol. modeling will be reported. In vivo tests of some compds. which interacted strongly with the central benzodiazepine receptors revealed reasonably potent anticonvulsant/anticonflict activity, and some of those which bind selectively to the peripheral-type (mitochondrial) benzodiazepine receptors are being examined as possible radiopharmaceuticals for imaging of tumors (and other disease states).
 IT 142074-27-9
 RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)
 (interaction with central and peripheral-type (mitochondrial) benzodiazepine receptors of imidazo[b]pyridazines in relation to anticonvulsant/anticonflict activity and activity as radiopharmaceuticals)
 RN 142074-27-9 CAPLUS
 CN Imidazo[1,2-a]pyrazine, 6-chloro-3-methoxy-2-(4-methylphenyl)- (9CI) (CA INDEX NAME)



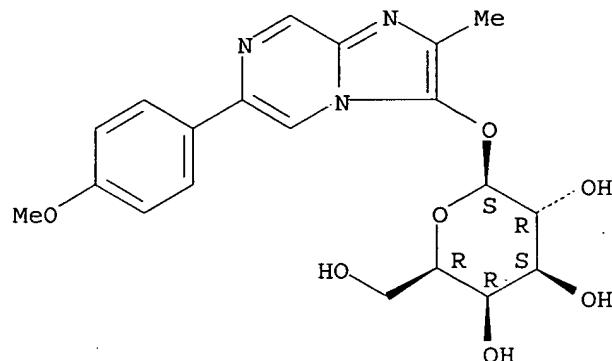
REFERENCE COUNT: 80 THERE ARE 80 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L24 ANSWER 10 OF 24 CAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 1997:48722 CAPLUS
 DOCUMENT NUMBER: 126:72331
 TITLE: Chemiluminescent substrate for enzyme immunoassay
 INVENTOR(S): Sakaki, Hidejiro; Mitani, Motohiro; Koinuma, Yasuyoshi; Totani, Yoshiaki
 PATENT ASSIGNEE(S): Nippon Oils & Fats Co Ltd, Japan
 SOURCE: Jpn. Kokai Tokkyo Koho, 6 pp.
 CODEN: JKXXAF

DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

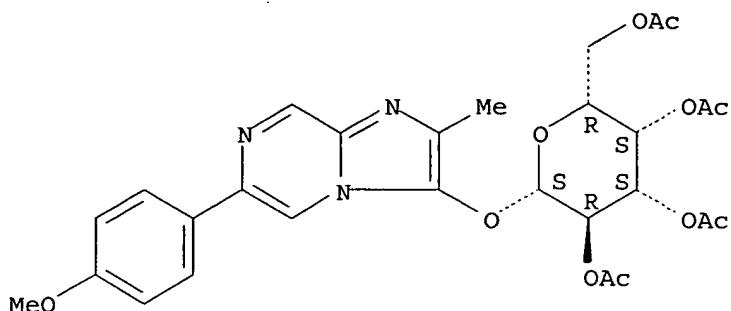
PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 08294397	A2	19961112	JP 1995-125617	19950427
PRIORITY APPLN. INFO.:	JP 1995-125617 19950427			
OTHER SOURCE(S):	MARPAT 126:72331			
AB	Chemiluminescent substrate for sugar-hydrolyzing enzyme is prepared for EIA. 3-(β -D-galactopyranosyloxy)-6-(4-methoxyphenyl)-2-methylimidazole[1,2-a]pyrazine was prepared from 6-(4-methoxyphenyl)-2-methyl-3-(tetra-O-acetyl- β -D-galactopyranosyloxy)imidazole[1,2-a]pyrazine, and used for chemiluminescent EIA.			
IT 159503-66-9P	RL: ARU (Analytical role, unclassified); SPN (Synthetic preparation); ANST (Analytical study); PREP (Preparation) (chemiluminescent substrate for EIA using carbohydrate-hydrolyzing enzyme)			
RN 159503-66-9 CAPLUS				
CN β -D-Galactopyranoside, 6-(4-methoxyphenyl)-2-methylimidazo[1,2-a]pyrazin-3-yl (9CI) (CA INDEX NAME)				

Absolute stereochemistry.

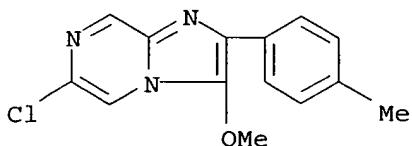


IT 177205-13-9P	RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent) (chemiluminescent substrate for EIA using carbohydrate-hydrolyzing enzyme)
RN 177205-13-9 CAPLUS	
CN β -D-Galactopyranoside, 6-(4-methoxyphenyl)-2-methylimidazo[1,2-a]pyrazin-3-yl, 2,3,4,6-tetraacetate (9CI) (CA INDEX NAME)	

Absolute stereochemistry.



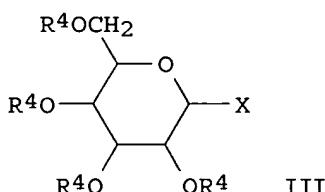
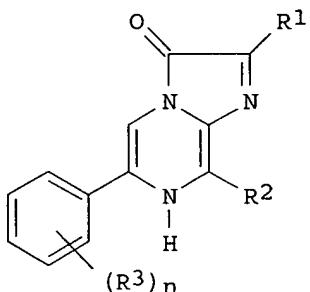
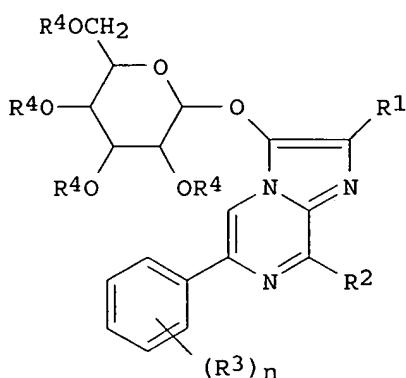
L24 ANSWER 11 OF 24 CAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 1996:397698 CAPLUS
 DOCUMENT NUMBER: 125:184868
 TITLE: Ligands for the central benzodiazepine receptor:
 structure-affinity relationship studies on
 imidazo[1,2-b]pyridazines
 AUTHOR(S): Matyus, Peter; Barlin, Gordon B.; Harrison, Peter W.;
 Wong, Margaret G.; Davies, Les P.
 CORPORATE SOURCE: Div. Neuroscience, Australian National Univ.,
 Canberra, 2601, Australia
 SOURCE: Australian Journal of Chemistry (1996), 49(4), 435-442
 CODEN: AJCHAS; ISSN: 0004-9425
 PUBLISHER: Commonwealth Scientific and Industrial Research
 Organization
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 AB Seventy-six imidazo[1,2-b]pyridazines and some bicyclic isomers have been
 analyzed and compared in terms of geometric and electronic requirements
 for binding to central benzodiazepine receptors. The binding sites
 identified for these compds. by mol. modeling are consistent with known
 benzodiazepine receptor-ligand interaction models. However, for the most
 active compds., addnl. binding sites are proposed.
 IT 142074-27-9
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological
 study, unclassified); PRP (Properties); BIOL (Biological study)
 (structure-affinity relationship of imidazopyridazines as ligands for
 the central benzodiazepine receptor)
 RN 142074-27-9 CAPLUS
 CN Imidazo[1,2-a]pyrazine, 6-chloro-3-methoxy-2-(4-methylphenyl)- (9CI) (CA
 INDEX NAME)



L24 ANSWER 12 OF 24 CAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 1996:335963 CAPLUS
 DOCUMENT NUMBER: 125:11354
 TITLE: Preparation of luciferin derivatives of *Umihotaru*
 (*Cypridina hilgendorfii*)

INVENTOR(S) : Mitani, Motohiro; Sakaki, Hidejiro; Koinuma, Yasuyoshi; Totani, Yoshiaki
 PATENT ASSIGNEE(S) : Nippon Oils & Fats Co., Ltd., Japan; NOF Corporation
 SOURCE: Jpn. Kokai Tokkyo Koho, 8 pp.
 CODEN: JKXXAF
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 08059686	A2	19960305	JP 1994-198770	19940823
JP 3648763	B2	20050518	JP 1994-198770	19940823
PRIORITY APPLN. INFO. :			CASREACT 125:11354; MARPAT 125:11354	
OTHER SOURCE(S) :				
GI				



- AB The title compds. (I; R1, R2 = H, C1-20 alkyl, C6-20 aryl, C7-19 arylalkyl; R3 = C1-5 alkyl or alkoxy; n = 0-5), which are useful as substrates for luminescent determination of sugar hydrolases such as α -D-galactosidase, are prepared by reacting imidazopyrazinone derivs. (II; R1 - R3, n = same as above) with sugar derivs. (III; X = halo; R4 = C1-7 acyl) in the presence of silver triflate and Na2HPO4. followed by solvolysis in the presence of an alkali. Thus, 0.1 g 6-(4-methoxyphenyl)-2-methylimidazo[1,2-a]pyrazin-3-one and 1.1 g Na2HPO4 were treated with 5 mL MeCN, 9 mL benzene, and 2.6 g mol. sieve 4A and stirred at room temperature for 1 h, treated with 0.18 g 2,3,4,6-tetra-O-acetyl- α -D-galactopyranosyl bromide and 0.37 g silver triflate, and stirred at room temperature for 2 h to give 39% 6-(4-methoxyphenyl)-2-methyl-3-(2,3,4,6-tetra-O-acetyl- α -D-galactopyranosyloxy)imidazo[1,2-a]pyrazine, which (0.5 g)

was treated with 3.5 mL MeOH and 1.8 mL concentrated aqueous NH₃ and stirred at 40° for 6 h 30 min to give 78% 6-(4-methoxyphenyl)-2-methyl-3-(α -D-galactopyranosyloxy)imidazo[1,2-a]pyrazine (IV). IV showed luminescence in the presence of β -D-galactosidase with correlation factor r = 0.992.

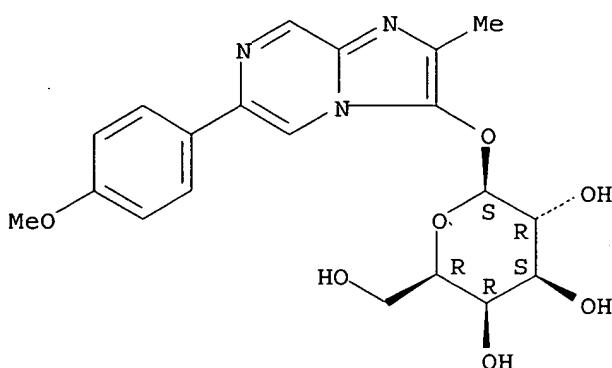
IT 159503-66-9P 177205-12-8P

RL: ARG (Analytical reagent use); SPN (Synthetic preparation); ANST (Analytical study); PREP (Preparation); USES (Uses)
(preparation of luciferin derivs. of Cypridina hilgendorfii as substrates for luminescent determination of sugar hydrolases)

RN 159503-66-9 CAPLUS

CN β -D-Galactopyranoside, 6-(4-methoxyphenyl)-2-methylimidazo[1,2-a]pyrazin-3-yl (9CI) (CA INDEX NAME)

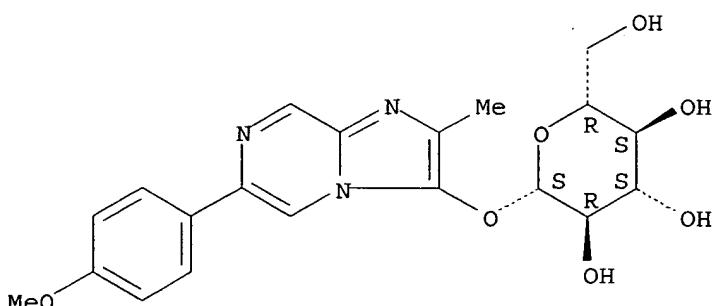
Absolute stereochemistry.



RN 177205-12-8 CAPLUS

CN β -D-Glucopyranoside, 6-(4-methoxyphenyl)-2-methylimidazo[1,2-a]pyrazin-3-yl (9CI) (CA INDEX NAME)

Absolute stereochemistry.



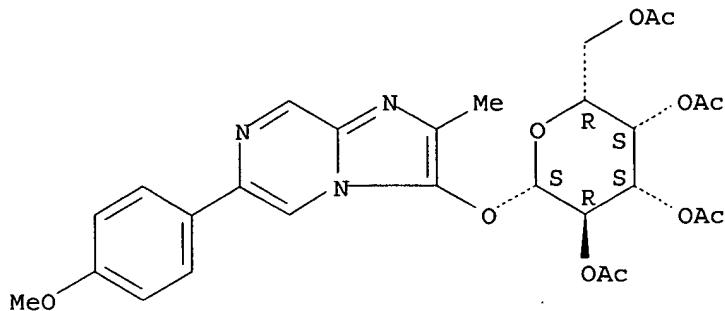
IT 177205-13-9P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
(preparation of luciferin derivs. of Cypridina hilgendorfii as substrates for luminescent determination of sugar hydrolases)

RN 177205-13-9 CAPLUS

CN β -D-Galactopyranoside, 6-(4-methoxyphenyl)-2-methylimidazo[1,2-a]pyrazin-3-yl, 2,3,4,6-tetraacetate (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L24 ANSWER 13 OF 24 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 1995:992122 CAPLUS

DOCUMENT NUMBER: 124:80192

TITLE: Enhancement effect of 2,6-O-dimethyl- β -cyclodextrin on the chemiluminescent detection of β -D-galactosidase using a Cypridina luciferin analog

AUTHOR(S): Mitani, Motohiro; Sakaki, Syujiro; Koinuma, Yasumi; Toya, Yoshiaki; Kosugi, Masanori

CORPORATE SOURCE: Tsukuba Res. Lab., NOF Corp., Tsukuba, 300-26, Japan

SOURCE: Analytical Sciences (1995), 11(6), 1013-15

CODEN: ANSCEN; ISSN: 0910-6340

PUBLISHER: Japan Society for Analytical Chemistry

DOCUMENT TYPE: Journal

LANGUAGE: English

AB β -Cyclodextrins enhanced the chemiluminescent detection of β -galactosidase using the Cypridina luciferin analog 3-(β -D-galactopyranosyloxy)-6-(4-methoxyphenyl)-2-methylimidazo[1,2-alpyrazine (β -Gal-MCLA) in the order 2,6-O-dimethyl- β -cyclodextrin > 2,3,6-O-trimethyl- β -cyclodextrin > β -cyclodextrin. Detection of mouse IgG by chemiluminescent enzyme immunoassay (CLEIA) using β -Gal-MCLA and β -galactosidase to amplify the signal was also enhanced by inclusion of 2,6-O-trimethyl- β -cyclodextrin.

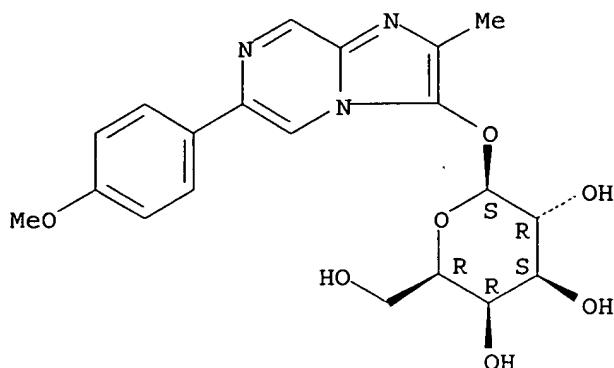
IT 159503-66-9

RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses) (enhancement effect of 2,6-O-dimethyl- β -cyclodextrin on the chemiluminescent detection of β -D-galactosidase using a Cypridina luciferin analog)

RN 159503-66-9 CAPLUS

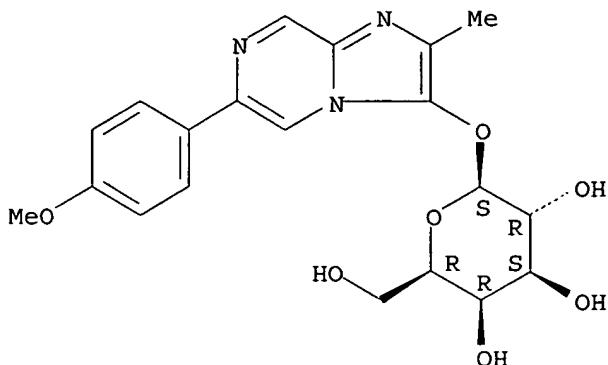
CN β -D-Galactopyranoside, 6-(4-methoxyphenyl)-2-methylimidazo[1,2-alpyrazin-3-yl (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L24 ANSWER 14 OF 24 CAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 1995:126975 CAPLUS
 DOCUMENT NUMBER: 122:4783
 TITLE: Chemiluminescent assay of β -D-galactosidase using Cypridina luciferin analog: 3-(β -D-galactopyranosyloxy)-6-(4-methoxyphenyl)-2-methylimidazo[1,2-a]pyrazine
 AUTHOR(S): Mitani, Motohiro; Sakaki, Syujiro; Koinuma, Yasumi; Toya, Yoshiaki; Kosugi, Masanori
 CORPORATE SOURCE: Tsukuba Res. Lab., NOF Corp., Ibaraki, 300-26, Japan
 SOURCE: Analytical Sciences (1994), 10(5), 813-14
 CODEN: ANSCEN; ISSN: 0910-6340
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 AB We prepared a new Cypridina luciferin analog, 3-(β -D-galactopyranosyloxy)-6-(4-methoxyphenyl)-2-methylimidazo[1,2-a]-pyrazine (β -Gal-MCLA) which can enzymically remove galactose to produce 2-methyl-6-(4-methoxyphenyl)-3,7-dihydroimidazo[1,2-a]pyrazine-3(7H)-one (MCLA), its autoxidn. follows, providing the chemiluminescence. β -Gal-MCLA was thus a useful chemiluminescent substrate for β -D-galactosidase determination
 IT 159503-66-9
 RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses) (chemiluminescent assay of β -D-galactosidase using Cypridina luciferin analog: 3-(β -D-galactopyranosyloxy)-6-(4-methoxyphenyl)-2-methylimidazo[1,2-a]pyrazine)
 RN 159503-66-9 CAPLUS
 CN β -D-Galactopyranoside, 6-(4-methoxyphenyl)-2-methylimidazo[1,2-a]pyrazin-3-yl (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L24 ANSWER 15 OF 24 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 1992:448498 CAPLUS

DOCUMENT NUMBER: 117:48498

TITLE: Imidazo[1,2-b]pyridazines. XII. Syntheses and central nervous system activities of some substituted imidazo[1,2-b]pyridazines and related imidazo[1,2-a]pyrimidines, imidazo[1,2-a]pyrimidines and imidazo[1,2-a]pyrazines

AUTHOR(S): Barlin, Gordon B.; Davies, Les P.; Ireland, Stephen J.; Ngu, Maria M. L.; Zhang, Jiankuo

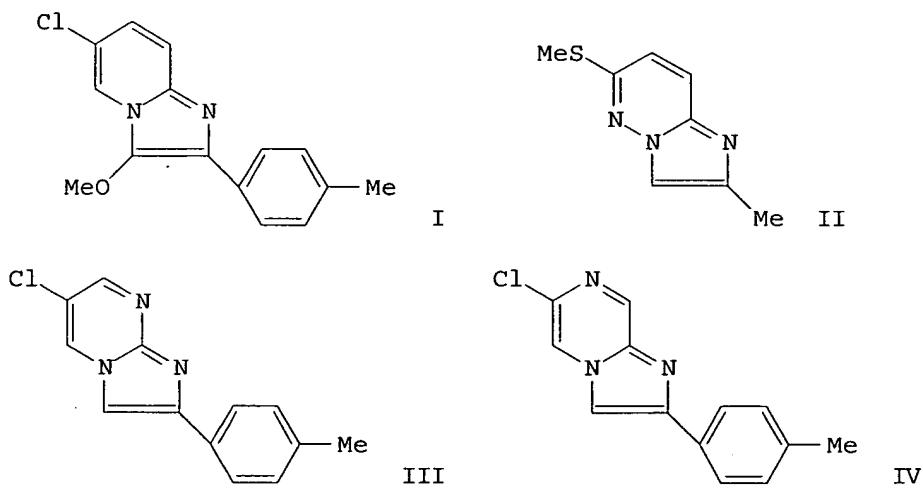
CORPORATE SOURCE: John Curtin Sch. Med. Res., Aust. Natl. Univ., Canberra, 2601, Australia

SOURCE: Australian Journal of Chemistry (1992), 45(5), 877-88
CODEN: AJCHAS; ISSN: 0004-9425

DOCUMENT TYPE: Journal

LANGUAGE: English

GI



AB Syntheses are reported for some 6-chloro(alkoxy, alkylthio and phenylthio)-3-benzamidomethyl(acetamidomethyl and methoxy)-2-arylimidazo[1,2-a]pyridines, e.g. I, and some corresponding imidazo[1,2-b]pyridazines, e.g. II, imidazo[1,2-a]pyrimidines, e.g. III,

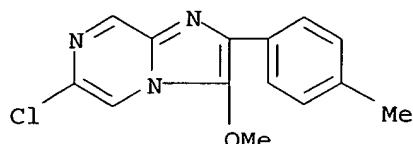
and imidazo[1,2-a]pyrazines, e.g. IV. Thus, 5-chloropyridin-2-amine was treated with p-tolylglyoxal to give I. IC₅₀ values (or percentage displacement) are reported and discussed for the displacement of [³H]diazepam from rat brain membrane by each of these compds. The imidazo[1,2-a]pyridines were generally slightly less potent than the imidazo[1,2-b]pyridazines but considerably more potent than the corresponding imidazo[1,2-a]pyrimidines or imidazo[1,2-a]pyrazines. Substitution of a 2-aryl group by a 2-alkyl group in imidazo[1,2-b]pyridazines led to significant loss of activity.

IT 142074-27-9P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation)
(preparation and central nervous system activity of)

RN 142074-27-9 CAPLUS

CN Imidazo[1,2-a]pyrazine, 6-chloro-3-methoxy-2-(4-methylphenyl)- (9CI) (CA INDEX NAME)



L24 ANSWER 16 OF 24 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 1987:153383 CAPLUS

DOCUMENT NUMBER: 106:153383

TITLE: Chemical studies of myctophina fish bioluminescence

AUTHOR(S): Inoue, Shoji; Okada, Kunisuke; Tanino, Hideo; Kakoi, Hisae

CORPORATE SOURCE: Fac. Pharm., Meijo Univ., Nagoya, 468, Japan

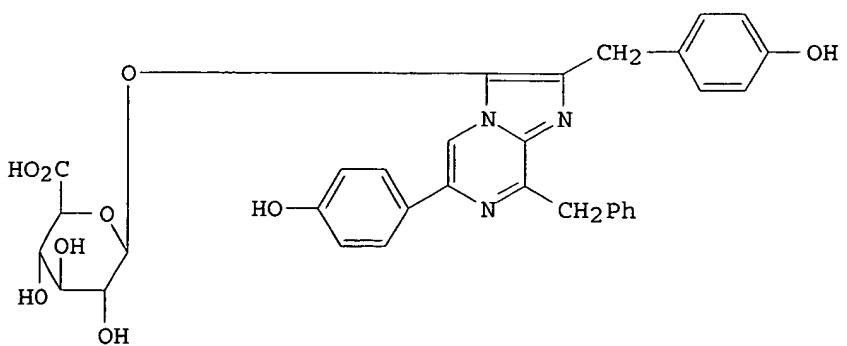
SOURCE: Chemistry Letters (1987), (2), 417-18

CODEN: CMLTAG; ISSN: 0366-7022

DOCUMENT TYPE: Journal

LANGUAGE: English

GI



I

AB A new type of masked watasenia preluciferin was isolated from the liver of

a myctophina fish (*Diaphus elucens*) and its structure was determined as watasenia preluciferyl β -D-glucopyranosiduronic acid (I).

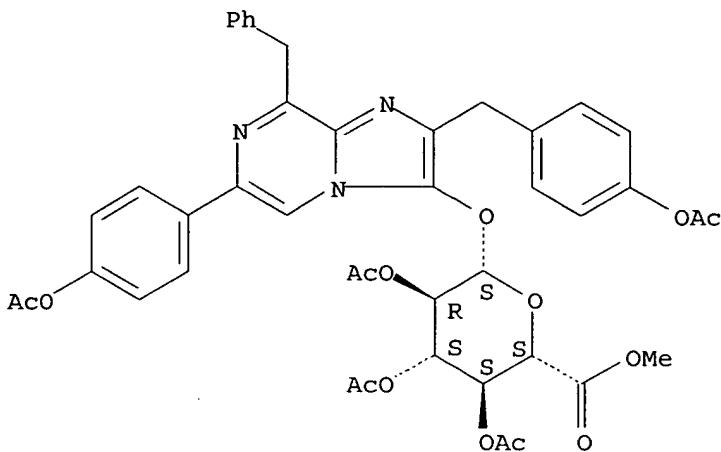
IT 107503-11-7

RL: RCT (Reactant); RACT (Reactant or reagent)
(deacetylation of)

RN 107503-11-7 CAPLUS

CN β -D-Glucopyranosiduronic acid, 6-[4-(acetoxy)phenyl]-2-[(4-(acetoxy)phenyl)methyl]-8-(phenylmethyl)imidazo[1,2-a]pyrazin-3-yl, methyl ester, 2,3,4-triacetate (9CI) (CA INDEX NAME)

Absolute stereochemistry.



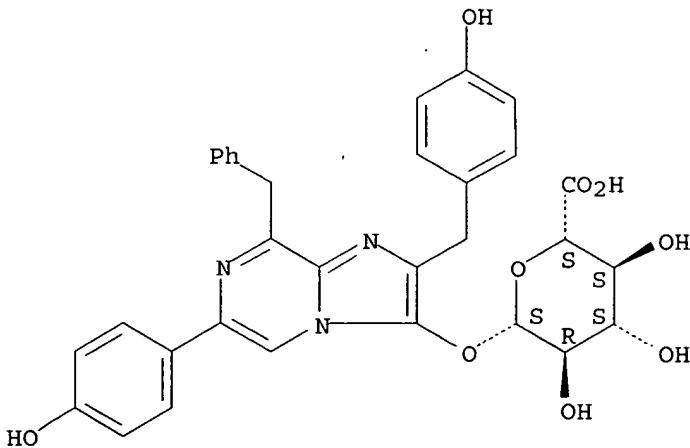
IT 107503-09-3

RL: BIOL (Biological study)
(of liver, of myctophina fish)

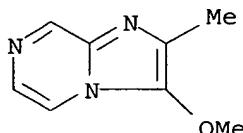
RN 107503-09-3 CAPLUS

CN β -D-Glucopyranosiduronic acid, 6-(4-hydroxyphenyl)-2-[(4-hydroxyphenyl)methyl]-8-(phenylmethyl)imidazo[1,2-a]pyrazin-3-yl (9CI)
(CA INDEX NAME)

Absolute stereochemistry. Rotation (-).



L24 ANSWER 17 OF 24 CAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 1985:55384 CAPLUS
 DOCUMENT NUMBER: 102:55384
 TITLE: Carbon-13 nuclear magnetic resonance spectra in the identification of N-, O- or S-methyl derivatives of some tautomeric hydroxy and mercapto nitrogen heterocycles
 AUTHOR(S): Barlin, Gordon B.; Brown, Desmond J.; Fenn, M. David
 CORPORATE SOURCE: John Curtin Sch. Med. Res., Aust. Natl. Univ., Canberra, 2601, Australia
 SOURCE: Australian Journal of Chemistry (1984), 37(11), 2391-5
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 AB Carbon-13 NMR spectroscopy, in contrast to 1H NMR spectroscopy, has been shown to provide a clear distinction in a variety of N heterocyclic systems between O-Me and nuclear N-Me groups. MeO groups occur in the range δ 53.20-61.87, nuclear N-Me groups at 34.29-49.62, and MeS groups at 12.35-14.55 for the compds. examined in CDCl₃. Data for N- and O-Me derivs. of pyridin-2 and -4-ol, the corresponding pyrimidines, and some S analogs were compared with those for the unmethylated parent compds.
 IT 87814-38-8
 RL: ANST (Analytical study)
 (identification of, carbon-13 NMR spectrometric)
 RN 87814-38-8 CAPLUS
 CN Imidazo[1,2-a]pyrazine, 3-methoxy-2-methyl- (9CI) (CA INDEX NAME)



L24 ANSWER 18 OF 24 CAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 1984:468065 CAPLUS
 DOCUMENT NUMBER: 101:68065
 TITLE: Mechanism of photoinactivation and re-activation in the bioluminescence system of the ctenophore Mnemiopsis
 AUTHOR(S): Anctil, Michel; Shimomura, Osamu
 CORPORATE SOURCE: Mar. Biol. Lab., Woods Hole, MA, 02543, USA
 SOURCE: Biochemical Journal (1984), 221(1), 269-72
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 AB The bioluminescence of *M. leidyi* takes place when the photoprotein mnemiopsis in the photocytes reacts with Ca²⁺. The luminescence is inhibited in sunlight and this photoinhibition is reversible by keeping the live specimens in the dark. Exts. of mnemiopsis are similarly photoinhibited, but the photoinhibition cannot be reversed in the dark. Photoinhibited mnemiopsis can be reactivated in the dark by incubation with coelenterazine and O₂ only in solns. having a pH very close to 9.0. The reactivation in vivo probably takes place in the same manner, using the coelenterazine that is supplied from its abundant storage form. Apparently, photoinactivation of mnemiopsis results in the dissociation of

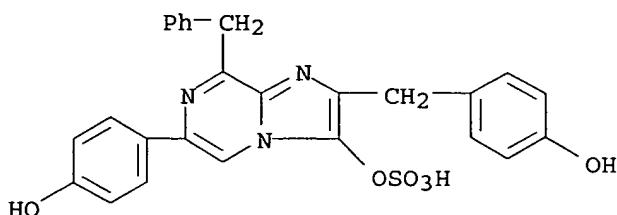
coelenterazine and O from the mol. of photoprotein; the dissociated form of the former mol. is an inactive form of coelenterazine, not free coelenterazine.

IT 65417-14-3

RL: BIOL (Biological study)
(of ctenophore)

RN 65417-14-3 CAPLUS

CN Imidazo[1,2-a]pyrazin-3-ol, 6-(4-hydroxyphenyl)-2-[(4-hydroxyphenyl)methyl]-8-(phenylmethyl)-, 3-(hydrogen sulfate), monosodium salt (9CI) (CA INDEX NAME)



● Na

L24 ANSWER 19 OF 24 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 1983:594926 CAPLUS

DOCUMENT NUMBER: 99:194926

TITLE: Imidazo[1,2-b]pyridazines and an imidazo[1,2-a]pyrazine from pyridazin- and pyrazinamines

AUTHOR(S): Barlin, Gordon B.; Brown, Desmond J.; Kadunc, Zdenka; Petric, Andrej; Stanovnik, Branka; Tisler, Miha

CORPORATE SOURCE: John Curtin Sch. Med. Res., Canberra, 2601, Australia

SOURCE: Australian Journal of Chemistry (1983), 36(6), 1215-20

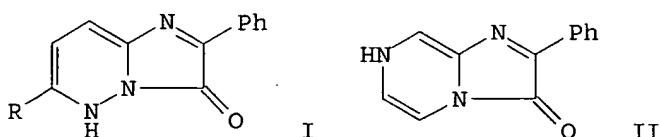
CODEN: AJCHAS; ISSN: 0004-9425

DOCUMENT TYPE: Journal

LANGUAGE: English

OTHER SOURCE(S): CASREACT 99:194926

GI

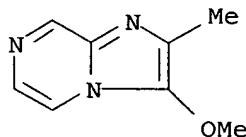


AB The ambiguous condensations of PhCOCHO with pyridazin-3-amines and pyrazin-2-amine give imidazopyridazinones I (R = H, Cl) and imidazopyrazinone II; resp. The former products exist as such, at least in the solid state, whereas the latter product exists to a large extent as the corresponding dipolar mol. The reactions, degradns., and NMR spectra of the products are discussed.

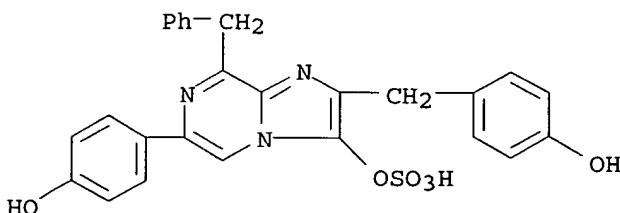
IT 87814-38-8P

RL: SPN (Synthetic preparation); PREP (Preparation)

(preparation of)
 RN 87814-38-8 CAPLUS
 CN Imidazo[1,2-a]pyrazine, 3-methoxy-2-methyl- (9CI) (CA INDEX NAME)



L24 ANSWER 20 OF 24 CAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 1980:107639 CAPLUS
 DOCUMENT NUMBER: 92:107639
 TITLE: Comparison of the amounts of key components in the bioluminescence systems of various coelenterates
 AUTHOR(S): Shimomura, Osamu; Johnson, Frank H.
 CORPORATE SOURCE: Dep. Biol., Princeton Univ., Princeton, NJ, 08540, USA
 SOURCE: Comparative Biochemistry and Physiology, Part B: Biochemistry & Molecular Biology (1979), 64B(1), 105-7
 CODEN: CBPBB8; ISSN: 0305-0491
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 AB Luciferase, photoprotein, free and protein-bound coelenterazine (I) and I enol-sulfate were assayed and compared in 5 bioluminescent coelenterates. Hydrozoans Aequorea aequorea and Halistaura cellularia contained photoprotein plus very small amts. of I enol-sulfate and luciferase activity, but no free I. Anthozoans Ptilosarcus gurneyi, Cavernularia obesa, and Renilla muelleri contained luciferase, I, and I enol-sulfate, but very little or no photoprotein. I existed mainly in a stabilized form bound to a Ca-binding protein. The bioluminescent reactions in the coelenterates were compared.
 IT 55779-47-0
 RL: BIOL (Biological study)
 (of coelenterates, bioluminescence in relation to)
 RN 55779-47-0 CAPLUS
 CN Imidazo[1,2-a]pyrazin-3-ol, 6-(4-hydroxyphenyl)-2-[(4-hydroxyphenyl)methyl]-8-(phenylmethyl)-, 3-(hydrogen sulfate) (9CI) (CA INDEX NAME)



L24 ANSWER 21 OF 24 CAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 1979:519858 CAPLUS
 DOCUMENT NUMBER: 91:119858
 TITLE: A Bioluminescence assay for PAP (3',5'-diphosphoadenosine) and PAPS (3'-phosphoadenylyl

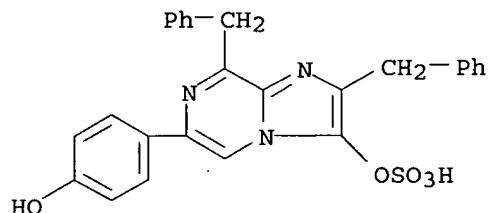
AUTHOR(S) : sulfate
 Anderson, James Michael; Hori, Kazuo; Cormier, Milton J.
 CORPORATE SOURCE: Boyd Grad. Stud. Res. Cent., Univ. Georgia, Athens, GA, 30602, USA
 SOURCE: Methods in Enzymology (1978), 57(Biolumin. Chemilumin.), 244-57
 CODEN: MENZAU; ISSN: 0076-6879
 DOCUMENT TYPE: Journal
 LANGUAGE: English

AB Procedures in the bioluminescence assay of PAP and PAPS using the luciferin-luciferase reaction in *Renilla reniformis* are described. The assay is sensitive to 0.1 pmol of PAP. The synthesis of the substrate benzyl luciferyl sulfate and isolation of luciferin sulfokinase and luciferase are also described.

IT 71369-28-3P
 RL: PREP (Preparation)
 (preparation of, as substrate for diphosphoadenosine and PAPS bioluminescence assay)

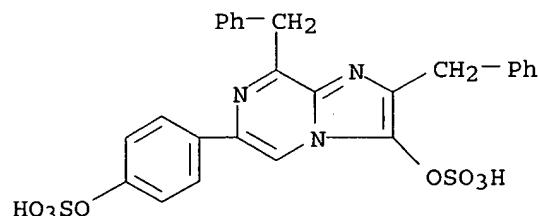
RN 71369-28-3 CAPLUS

CN Imidazo[1,2-a]pyrazin-3-ol, 6-(4-hydroxyphenyl)-2,8-bis(phenylmethyl)-, 3-(hydrogen sulfate), monopotassium salt (9CI) (CA INDEX NAME)



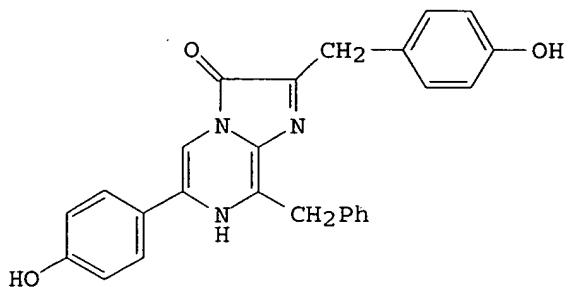
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IT 71369-27-2
 RL: RCT (Reactant); RACT (Reactant or reagent)
 (reaction of, with aryl sulfatase)
 RN 71369-27-2 CAPLUS
 CN Imidazo[1,2-a]pyrazin-3-ol, 2,8-bis(phenylmethyl)-6-[4-(sulfoxy)phenyl]-, hydrogen sulfate (ester), dipotassium salt (9CI) (CA INDEX NAME)

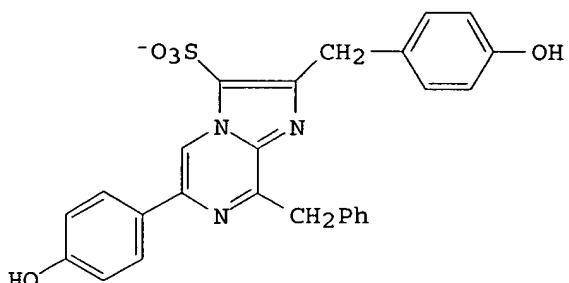


● 2 K

L24 ANSWER 22 OF 24 CAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 1978:50764 CAPLUS
 DOCUMENT NUMBER: 88:50764
 TITLE: Complete structure of Renilla luciferin and luciferyl sulfate
 AUTHOR(S): Inoue, Shoji; Kakoi, Hisae; Murata, Mikiko; Goto, Toshio; Shimomura, Osamu
 CORPORATE SOURCE: Fac. Pharm., Meijo Univ., Nagoya, Japan
 SOURCE: Tetrahedron Letters (1977), (31), 2685-8
 CODEN: TELEAY; ISSN: 0040-4039
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 GI

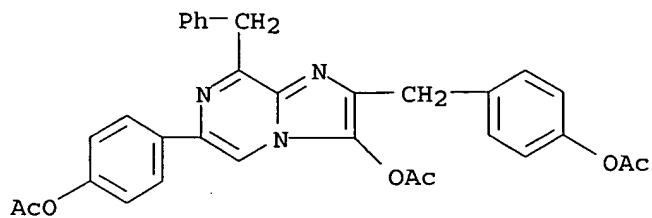


I



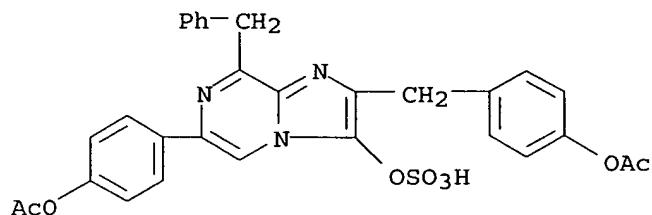
II

- AB Examination of *Renilla* exts. showed that *Renilla luciferin* is coelenterazine (I). The structure of natural luciferyl sulfate was determined as II by comparison of natural and synthetic II. II was synthesized from I by sequential treatment with (AcO)₂O, MeOH/NH₃, and pyridine-SO₃ complex and hydrolysis with MeOH/NaOH.
- IT 65417-16-5P 65417-17-6P
 RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
 (preparation and hydrolysis of)
- RN 65417-16-5 CAPLUS
- CN Imidazo[1,2-a]pyrazin-3-ol, 6-[4-(acetyloxy)phenyl]-2-[[4-(acetyloxy)phenyl]methyl]-8-(phenylmethyl)-, acetate (ester) (9CI) (CA INDEX NAME)



RN 65417-17-6 CAPLUS

CN Imidazo[1,2-a]pyrazin-3-ol, 6-[4-(acetyloxy)phenyl]-2-[[4-(acetyloxy)phenyl]methyl]-8-(phenylmethyl)-, hydrogen sulfate (ester), sodium salt (9CI) (CA INDEX NAME)



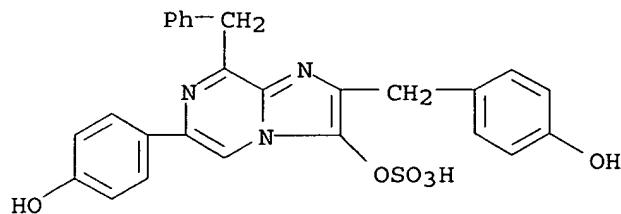
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IT 65417-14-3P

RL: PRP (Properties); SPN (Synthetic preparation); PREP (Preparation)
(preparation and structure of)

RN 65417-14-3 CAPLUS

CN Imidazo[1,2-a]pyrazin-3-ol, 6-(4-hydroxyphenyl)-2-[(4-hydroxyphenyl)methyl]-8-(phenylmethyl)-, 3-(hydrogen sulfate), monosodium salt (9CI) (CA INDEX NAME)



● Na

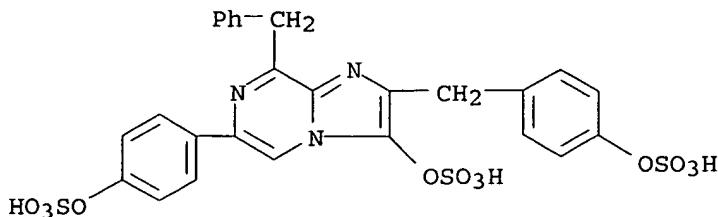
IT 65417-15-4P

RL: SPN (Synthetic preparation); PREP (Preparation)
(preparation of)

RN 65417-15-4 CAPLUS

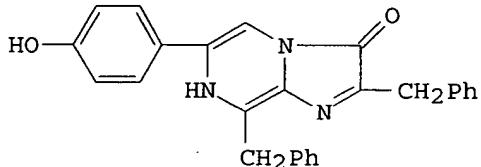
CN Imidazo[1,2-a]pyrazin-3-ol, 8-(phenylmethyl)-6-[4-(sulfooxy)phenyl]-2-[[4-(sulfooxy)phenyl]methyl]-, hydrogen sulfate (ester), trisodium salt (9CI)

(CA INDEX NAME)



●3 Na

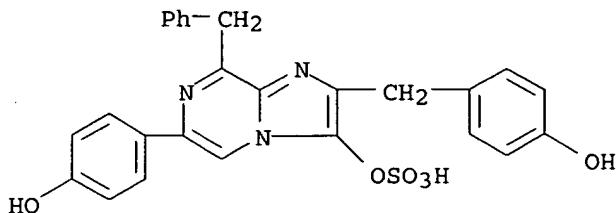
L24 ANSWER 23 OF 24 CAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 1977:596264 CAPLUS
 DOCUMENT NUMBER: 87:196264
 TITLE: Substrate and substrate analog binding properties of
 Renilla luciferase
 AUTHOR(S): Matthews, John C.; Hori, Kazuo; Cormier, Milton J.
 CORPORATE SOURCE: Dep. Biochem., Univ. Georgia, Athens, GA, USA
 SOURCE: Biochemistry (1977), 16(24), 5217-20
 CODEN: BICHAW; ISSN: 0006-2960
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 GI



I

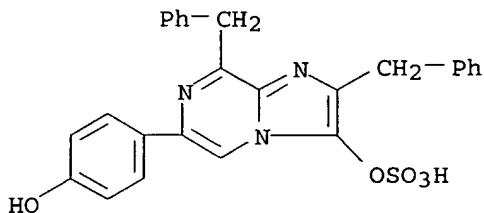
- AB The binding characteristics of luciferin, luciferin analogs (e.g. I), and competitive inhibitors of the luciferin-luciferase reaction were studied. Luciferin binding and orientation in the single luciferin binding site of luciferase from *R. reniformis* are highly specific for and dependent upon the 3 group substituents of the luciferin mol., whereas the imidazolone-pyrazine nucleus of luciferin is not directly involved in binding. Anaerobic luciferin binding promotes a rapid concentration-dependent aggregation of luciferase which results in irreversible inactivation of the enzyme. This aggregation phenomenon is not observed upon binding of oxyluciferin, luciferyl sulfate, or luciferin analogs in which the substituent at the 2 position of the imidazolone-pyrazine ring has been substantially altered.
- IT 55779-47-0 64750-83-0
 RL: PROC (Process)
 (luciferase binding of, structural factors in)
- RN 55779-47-0 CAPLUS
- CN Imidazo[1,2-a]pyrazin-3-ol, 6-(4-hydroxyphenyl)-2-[(4-hydroxyphenyl)methyl]-8-(phenylmethyl)-, 3-(hydrogen sulfate) (9CI) (CA

(INDEX NAME)



RN 64750-83-0 CAPLUS

CN Imidazo[1,2-a]pyrazin-3-ol, 6-(4-hydroxyphenyl)-2,8-bis(phenylmethyl)-, 3-(hydrogen sulfate) (9CI) (CA INDEX NAME)



L24 ANSWER 24 OF 24 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 1975:405600 CAPLUS

DOCUMENT NUMBER: 83:5600

TITLE: Chemical nature of bioluminescence systems in coelenterates

AUTHOR(S): Shimomura, Osamu; Johnson, Frank H.

CORPORATE SOURCE: Dep. Biol., Princeton Univ., Princeton, NJ, USA

SOURCE: Proceedings of the National Academy of Sciences of the United States of America (1975), 72(4), 1546-9

CODEN: PNASA6; ISSN: 0027-8424

DOCUMENT TYPE: Journal

LANGUAGE: English

AB Anal. of substances involved in light-emitting reactions among bioluminescent coelenterates revealed a pronounced uniformity in the structural features of initial reactants, i.e., luciferins and photoprotein chromophores, as well as the light-emitter product. This product is structurally identical among the different classes of coelenterates; i.e., Hydrozoa (the jellyfish, Aequorea), Anthozoa (the sea cactus, Cavernularia; sea pansy, Renilla; and sea pen, Leioptilus), and very likely also the Scyphozoa (the jellyfish, Pelagia). In each of these instances the reaction product, 2-(p-hydroxyphenylacetyl)amino-3-benzyl-5-(p-hydroxyphenyl) pyrazine, is the actual light-emitter, whether it occurs in a Ca²⁺-triggered photoprotein type of luminescence or in a luciferin-luciferase type. The evidence indicates that in certain coelenterates, e.g., Cavernularia, these 2 types are equally significant, whereas in others (Renilla and Leioptilus) the luciferin-luciferase type predominates over the Ca-triggerable photoprotein type. Only the photoprotein type functions in the luciferaseless jellyfish, Aequorea. In all instances investigated, the structure of the light-emitter prior to the luminescence reaction appears to be essentially the same as that of

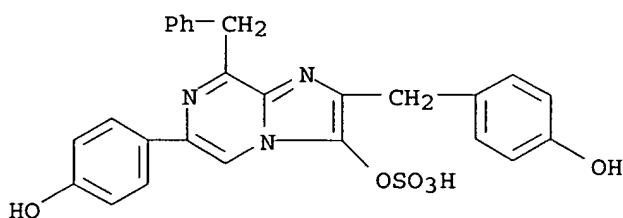
the chromophore of unreacted aequorin. The product of the luminescence reaction is absent in exts. of nonluminous species. However, a product very similar to that of luminescent coelenterates occurs also in representatives of other phyla, including the cephalopod molluscs, e.g., the "firefly squid" Watasenia and probably various ctenophores as well.

IT 55779-47-0

RL: BIOL (Biological study)
(in calcium-induced luminescence of coelenterates)

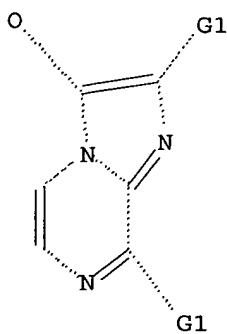
RN 55779-47-0 CAPLUS

CN Imidazo[1,2-a]pyrazin-3-ol, 6-(4-hydroxyphenyl)-2-[(4-hydroxyphenyl)methyl]-8-(phenylmethyl)-, 3-(hydrogen sulfate) (9CI) (CA INDEX NAME)



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L1 STR

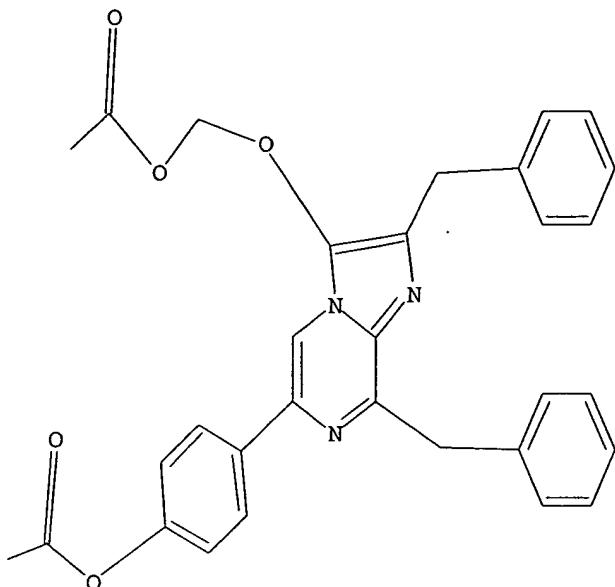
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L5

STR



Structure attributes must be viewed using STN Express query preparation.

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 L13 1 SEA FILE=CAPLUS ABB=ON PLU=ON L12

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L13 ANSWER 1 OF 1 CAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 2003:376823 CAPLUS
 DOCUMENT NUMBER: 138:365147
 TITLE: Compositions, methods and kits pertaining to luminescent compounds
 INVENTOR(S): Wood, Keith; Hawkins, Erika; Scurria, Mike; Klaubert, Dieter
 PATENT ASSIGNEE(S): Promega Corporation, USA
 SOURCE: PCT Int. Appl., 60 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2003040100	A1	20030515	WO 2002-US34972	20021101
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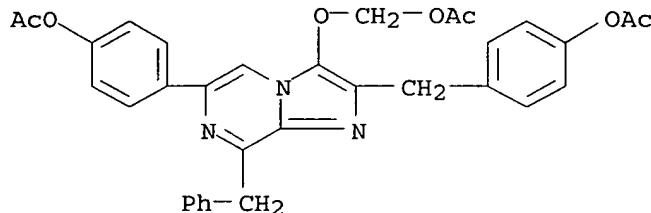
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 US 2003153090 A1 20030814 US 2001-53482 20011102
 CA 2462506 AA 20030515 CA 2002-2462506 20021101
 EP 1451155 A1 20040901 EP 2002-802815 20021101
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 JP 2005515977 T2 20050602 JP 2003-542146 20021101
 PRIORITY APPLN. INFO.: US 2001-53482 A 20011102
 WO 2002-US34972 W 20021101

OTHER SOURCE(S): MARPAT 138:365147

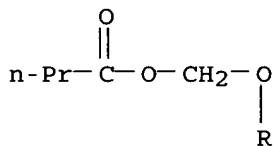
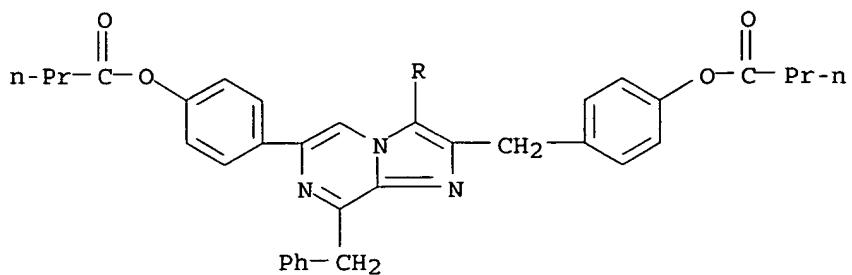
AB A method of measuring the enzymic activity of a luciferase includes contacting a luminogenic protein, such as a luciferase, with a protected luminophore to form a composition; and detecting light produced from the composition

The protected luminophore provides increased stability and improved signal-to-background ratios relative to the corresponding unmodified coelenterazine.

IT 524066-92-0P 524066-93-1P 524066-94-2P
 RL: ARU (Analytical role, unclassified); SPN (Synthetic preparation); ANST (Analytical study); PREP (Preparation)
 (compns., methods and kits pertaining to luminescent compds.)
 RN 524066-92-0 CAPLUS
 CN Phenol, 4-[3-[(acetyloxy)methoxy]-2-[[4-(acetyloxy)phenyl]methyl]-8-(phenylmethyl)imidazo[1,2-a]pyrazin-6-yl], acetate (ester) (9CI) (CA INDEX NAME)

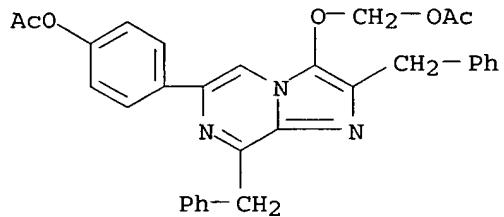


RN 524066-93-1 CAPLUS
 CN Butanoic acid, 4-[3-[(1-oxobutoxy)methoxy]-2-[[4-(1-oxobutoxy)phenyl]methyl]-8-(phenylmethyl)imidazo[1,2-a]pyrazin-6-yl]phenyl ester (9CI) (CA INDEX NAME)



RN 524066-94-2 CAPLUS

CN Phenol, 4-[3-[(acetyloxy)methoxy]-2,8-bis(phenylmethyl)imidazo[1,2-a]pyrazin-6-yl]-, acetate (ester) (9CI) (CA INDEX NAME)



REFERENCE COUNT:

8

THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT